- key terms

#### 10/601844

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9 APR 2006 HIGHEST RN 879846-78-3 STRUCTURE FILE UPDATES: 9 APR 2006 HIGHEST RN 879846-78-3 DICTIONARY FILE UPDATES:

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TSCA INFORMATION NOW CURRENT THROUGH January 6, 2006

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\*\*\*\*\*\*\*\*\*\*\*\*\*\*\*\*\*\*\* \* The CA roles and document type information have been removed from \* \* the IDE default display format and the ED field has been added, \* effective March 20, 2005. A new display format, IDERL, is now \* available and contains the CA role and document type information. \* \*\*\*\*\*\*\*\*\*\*\*\*\*\*\*\*\*

Structure search iteration limits have been increased. See HELP SLIMITS for details.

REGISTRY includes numerically searchable data for experimental and predicted properties as well as tags indicating availability of experimental property data in the original document. For information on property searching in REGISTRY, refer to:

http://www.cas.org/ONLINE/UG/regprops.html

E LEVOCETIRIZINE/CN 5 L1 2 S E3-4

ANSWER 1 OF 2 REGISTRY COPYRIGHT 2006 ACS on STN L1

130018-87-0 REGISTRY RN

ED Entered STN: 26 Oct 1990

Acetic acid, [2-[4-[(R)-(4-chlorophenyl)phenylmethyl]-1-CN piperazinyl]ethoxy]-, dihydrochloride (9CI) (CA INDEX NAME) OTHER CA INDEX NAMES:

Acetic acid, [2-[4-[(4-chlorophenyl)phenylmethyl]-1piperazinyl]ethoxy]-, dihydrochloride, (R)-

OTHER NAMES:

CN Levocetirizine dihydrochloride

CN Xusal

FS STEREOSEARCH

MF C21 H25 Cl N2 O3 . 2 Cl H

SR

STN Files: BIOSIS, CA, CAPLUS, CHEMCATS, IMSPATENTS, IMSRESEARCH, LC MRCK\*, PATDPASPC, PS, TOXCENTER, USPAT2, USPATFULL (\*File contains numerically searchable property data)

CRN (130018-77-8)

Absolute stereochemistry. Rotation (+).

## •2 HCl

17 REFERENCES IN FILE CA (1907 TO DATE)
17 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L1 ANSWER 2 OF 2 REGISTRY COPYRIGHT 2006 ACS on STN

RN 130018-77-8 REGISTRY

ED Entered STN: 26 Oct 1990

CN Acetic acid, [2-[4-[(R)-(4-chlorophenyl)phenylmethyl]-1-piperazinyl]ethoxy]- (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Acetic acid, [2-[4-[(4-chlorophenyl)phenylmethyl]-1-piperazinyl]ethoxy]-, (R)-

OTHER NAMES:

CN (-)-Cetirizine

CN Levocetirizine

CN Xyzal

FS STEREOSEARCH

DR 744169-44-6

MF C21 H25 C1 N2 O3

CI COM

SR CA

LC STN Files: ADISNEWS, BIOSIS, BIOTECHNO, CA, CAPLUS, CHEMCATS, CIN, EMBASE, IMSPATENTS, IMSRESEARCH, IPA, MEDLINE, MRCK\*, PATDPASPC, PHAR, PROMT, PROUSDDR, PS, RTECS\*, SYNTHLINE, TOXCENTER, USAN, USPAT2, USPATFULL

(\*File contains numerically searchable property data)

Absolute stereochemistry. Rotation (+).

\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

102 REFERENCES IN FILE CA (1907 TO DATE)
2 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
102 REFERENCES IN FILE CAPLUS (1907 TO DATE)

#### E CETIRIZINE DIHYDROCHLORIDE/CN 2 S E2-3 L2 ANSWER 1 OF 2 REGISTRY COPYRIGHT 2006 ACS on STN L2 83881-52-1 REGISTRY RN Entered STN: 16 Nov 1984 ED Acetic acid, [2-[4-[(4-chlorophenyl)phenylmethyl]-1-CN piperazinyl]ethoxy]-, dihydrochloride (9CI) (CA INDEX NAME) OTHER NAMES: Alercet CN CN Alerid CN Alerlisin CN Cesta CN Cetirizine dihydrochloride CN Cetirizine hydrochloride CN Cetrine Cetrizet CN CN Cistamine CN Formistin Histazine CN CN Nosemin P 071 CN CN Reactine CN Riztec CN Ryzen CN Sancotec CN Selitex CN Triz UCB-P 071 CN CN Virlix CN Zeran CN Zirtec CN Zirtek CN Zirtin CN Zyrlex CN Zyrtec Zyrzine CN DR 130018-82-5 MF C21 H25 Cl N2 O3 . 2 Cl H CI LC STN Files: ADISNEWS, ANABSTR, BIOSIS, BIOTECHNO, CA, CAPLUS, CASREACT, CBNB, CHEMCATS, CIN, CSCHEM, DIOGENES, EMBASE, IMSPATENTS, IMSRESEARCH, IPA, MEDLINE, MRCK\*, PHAR, PROMT, PS, RTECS\*, SCISEARCH, SYNTHLINE, TOXCENTER, USAN, USPAT2, USPATFULL (\*File contains numerically searchable property data) CRN (83881-51-0)

#### ●2 HCl

\*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

177 REFERENCES IN FILE CA (1907 TO DATE)
177 REFERENCES IN FILE CAPLUS (1907 TO DATE)

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FILE COVERS 1907 - 10 Apr 2006 VOL 144 ISS 16 FILE LAST UPDATED: 9 Apr 2006 (20060409/ED)

XUSAL OR XYZAL

Effective October 17, 2005, revised CAS Information Use Policies apply. They are available for your review at:

http://www.cas.org/infopolicy.html

L1	2	SEA FILE=REGISTRY ABB=ON PLU=ON (LEVOCETIRIZINE/CN OR
		"LEVOCETIRIZINE DIHYDROCHLORIDE"/CN)
L3	121	SEA FILE=CAPLUS ABB=ON PLU=ON L1 OR LEVOCETIRIZINE OR
		XUSAL OR XYZAL
L4	1502	SEA FILE=CAPLUS ABB=ON PLU=ON (CHLOROPHENYL? OR (CL OR
		CHLORO) (W) (PH OR PHENYL?)) (S) ACETIC
L5		SEA FILE=CAPLUS ABB=ON PLU=ON L4(S)PIPERAZIN?
L6	80	SEA FILE=CAPLUS ABB=ON PLU=ON (L3 OR L5) AND (THERAP? OR
		TREAT? OR PREVENT? OR PROPHYLACT? OR PROPHYLAX?)
L7	3	SEA FILE=CAPLUS ABB=ON PLU=ON L6 AND (EXCIPIENT OR
		(STABILIS? OR STABILIZ? OR SUSPEND? OR SUSPENS?) (5A) AGENT)
L1	2	SEA FILE=REGISTRY ABB=ON PLU=ON (LEVOCETIRIZINE/CN OR
		"LEVOCETIRIZINE DIHYDROCHLORIDE"/CN)
L3	121	SEA FILE=CAPLUS ABB=ON PLU=ON L1 OR LEVOCETIRIZINE OR

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L4
            1502 SEA FILE=CAPLUS ABB=ON PLU=ON (CHLOROPHENYL? OR (CL OR
                  CHLORO) (W) (PH OR PHENYL?)) (S) ACETIC
              31 SEA FILE=CAPLUS ABB=ON PLU=ON L4(S)PIPERAZIN?
L5
              80 SEA FILE=CAPLUS ABB=ON PLU=ON (L3 OR L5) AND (THERAP? OR
Lб
                  TREAT? OR PREVENT? OR PROPHYLACT? OR PROPHYLAX?)
              19 SEA FILE=CAPLUS ABB=ON PLU=ON L6 AND (ORAL? OR TABLET OR
L8
                  PILL OR CAPSUL? OR PER OS OR MOUTH)
L9
              19 S L7 OR L8
     ANSWER 1 OF 19 CAPLUS COPYRIGHT 2006 ACS on STN
L9
     Entered STN: 28 Oct 2005
                            2005:1154350 CAPLUS
ACCESSION NUMBER:
DOCUMENT NUMBER:
                            143:411084
TITLE:
                            Pharmaceutical composition for treating
                            hair loss and benign prostatic hyperplasia
                            Lee, Eun-Joo
INVENTOR(S):
PATENT ASSIGNEE(S):
                            Lee, Eun-Joo, S. Korea
SOURCE:
                            PCT Int. Appl., 36 pp.
                            CODEN: PIXXD2
DOCUMENT TYPE:
                            Patent
                            English
LANGUAGE:
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
     PATENT NO.
                            KIND
                                    DATE
                                                APPLICATION NO.
                                                                          DATE
                                              WO 2005-KR1063 20050413
                            A1
     WO 2005099653
                                    20051027
          W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA,
              CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI,
              GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KP, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, IZ, VC, VN, VII, 70, 7M, 7M, 7M
              UZ, VC, VN, YU, ZA, ZM, ZW
          RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW,
              AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ,
              DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, MC,
              NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA,
              GN, GQ, GW, ML, MR, NE, SN, TD, TG
                                                 KR 2004-25869
PRIORITY APPLN. INFO.:
                                                                        A 20040414
                                                  KR 2005-29495
                                                                        A 20050408
```

The present invention is related to an orally administered pharmaceutical composition for the prevention of hair loss; having the effects of hair toning, and growth; and for the treatment of female hirsutism and benign prostatic hyperplasia. The composition is nonsteroidal in nature and is advantageous in that it has no side effects such as lowering of sexual function, shown in the conventional oral treatment agents of related diseases. The average weight of prostate glands of the comparative group to which finasteride is administered, or of the exptl. group to which a pharmaceutical composition having 2-[2-[4-[(4-chlorophenyl))phenylmethyl]-1-piperazinyl]ethoxy]acetic acid dihydrochloride after inducing enlargement of prostate glands with TP shows an almost similar value to the weight of prostate glands of the solvent control group to which only vehicles are administered., which

means that the enlarged size of prostate glands is recovered to the normal size and there are assured effects of treatment.

THERE ARE 4 CITED REFERENCES AVAILABLE FOR REFERENCE COUNT: THIS RECORD. ALL CITATIONS AVAILABLE IN THE

RE FORMAT

ANSWER 2 OF 19 CAPLUS COPYRIGHT 2006 ACS on STN

Entered STN: 22 Sep 2005

2005:1017930 CAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER: 143:278249

New oral antihistamines in pediatrics TITLE:

and safety of antihistamines in children

Ones, Ulker; Tamay, Zeynep AUTHOR(S):

Medical Faculty, Department of Pediatrics, CORPORATE SOURCE:

Division of Allergy and Chest Diseases, Istanbul

University, Istanbul, Turk.

Current Medicinal Chemistry: Anti-Inflammatory & SOURCE:

Anti-Allergy Agents (2005), 4(5), 495-506

CODEN: CMCAGM; ISSN: 1568-0142

Bentham Science Publishers Ltd. PUBLISHER:

Journal; General Review DOCUMENT TYPE:

English LANGUAGE:

A review. H1 antihistamines are first line drugs in the treatment of allergic rhinitis and chronic idiopathic urticaria and widely used in children as well as in adults. Although first-generation antihistamines are effective in relieving allergic symptoms, they are not preferred because of their sedative side effects. The earliest "second generation" antihistamines, terfenadine and astemizole, non-sedating alternatives to the first generation counterparts are not commonly used due to their potential arythmogenic effects. The newer second-generation antihistamines such as loratadine, fexofenadine, mizolastine, ebastine, cetirizine, levocetirizine and desloratadine have been shown to be efficacious and well tolerated with addnl. anti-inflammatory effects and lacking cardiotoxic potential activity in adults. The early treatment of atopic children study, the long term clin. trial with cetirizine of infants with atopic dermatitis demonstrated that cetirizine delayed the onset of asthma in patients sensitized to grass pollen or house dust mite; and also reduced the duration and the amount of topical steroids used in the treatment of atopic dermatitis. In the Preventia I study, which was designed to evaluate the efficacy of loratadine in reducing the number of respiratory infections in young children at risk of recurrent infections, loratadine was not significantly different from placebo. Both drugs were found to have a similar safety profile to that of placebo confirming their long-term use in infants and children. Pediatric formulation of desloratadine, which has favorable effect on nasal congestion, is marketed worldwide now. The effectiveness of new antihistamines in the treatment of urticaria in pediatric age group is based on extrapolation of adult studies performed in this area. Further studies with new antihistamines are needed for their evidence-based use in children with urticaria and atopic dermatitis.

ANSWER 3 OF 19 CAPLUS COPYRIGHT 2006 ACS on STN L9

152

Entered STN: 25 Aug 2005

REFERENCE COUNT:

2005:888933 CAPLUS ACCESSION NUMBER:

> Searcher : Shears 571-272-2528

RE FORMAT

THERE ARE 152 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE

DOCUMENT NUMBER: 143:206440

TITLE: Use of levocetirizine for the

preparation of a drug

INVENTOR(S):
Kouzan, Serge

PATENT ASSIGNEE(S): UCB Farchim SA, Switz. SOURCE: PCT Int. Appl., 13 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PAT	PATENT NO.						KIND DATE				APPLICATION NO.					
						-							<b>-</b> -			
WO	2005	0773	71		A1		2005	0825	Ī	WO 2	005-1	EP50	543		2	0050208
	W:	ΑE,	ΑG,	AL,	AM,	AT,	ΑU,	ΑZ,	BA,	BB,	BG,	BR,	BW,	BY,	BZ,	CA,
		CH,	CN,	CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	EG,	ES,	FI,
		GB,	GD,	GE,	GH,	GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	ΚE,	KG,	KP,
		KR,	ΚZ,	LC,	LK,	LR,	LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,
		MX,	MZ,	NA,	NI,	NO,	ΝZ,	OM,	PG,	PH,	PL,	PT,	RO,	RU,	SC,	SD,
		SE,	SG,	SK,	SL,	SY,	ТJ,	TM,	TN,	TR,	TT,	TZ,	UA,	UG,	US,	UZ,
		VC,	VN,	YU,	ZA,	ZM,	ZW									
	RW:	BW,	GH,	GM,	ΚE,	LS,	MW,	MZ,	NA,	SD,	SL,	SZ,	ΤŻ,	ŪG,	ZM,	ZW,
		AM,	ΑZ,	BY,	KG,	ΚZ,	MD,	RU,	ТJ,	TM,	ΑT,	ΒE,	ВG,	CH,	CY,	CZ,
		DE,	DK,	EE,	ES,	FI,	FR,	GB,	GR,	ΗU,	ΙE,	IS,	IT,	LT,	LU,	MC,
		NL,	PL,	PT,	RO,	SE,	SI,	SK,	TR,	BF,	ВJ,	CF,	CG,	CI,	CM,	GA,
		GN,	GQ,	GW,	ML,	MR,	ΝE,	SN,	-							
PRIORITY	APP	LN.	INFO	.:					:	EP 2	004-	3109		1	A 2	0040212

AB The present invention relates to a pharmaceutical use of levocetirizine for the prevention of symptoms or exacerbation of allergic asthma.

IT 130018-77-8, Levocetirizine 130018-87-0,

Levocetirizine dihydrochloride

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(levocetirizine for the preparation of a drug to treat

allergic asthma)

REFERENCE COUNT: 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR

THIS RECORD. ALL CITATIONS AVAILABLE IN THE

RE FORMAT

L9 ANSWER 4 OF 19 CAPLUS COPYRIGHT 2006 ACS on STN

ED Entered STN: 09 Aug 2005

ACCESSION NUMBER: 2005:704901 CAPLUS

DOCUMENT NUMBER: 143:472299

TITLE: Levocetirizine: Pharmacokinetics and

pharmacodynamics in children age 6 to 11 years

AUTHOR(S): Simons, F. Estelle R.; Simons, Keith J.

CORPORATE SOURCE: Department of Pediatrics and Child Health,

Department of Immunology, Canadian Institutes of Health Research National Training Program in Allergy and Asthma, Fac. Med., Univ. Manitoba,

Winnipeg, MB, Can.

SOURCE: Journal of Allergy and Clinical Immunology (2005),

116(2), 355-361

CODEN: JACIBY; ISSN: 0091-6749

PUBLISHER: Elsevier DOCUMENT TYPE: Journal

• • • • • • • • •

LANGUAGE: English

The pharmacokinetics and pharmacodynamics of medications may differ between children and adults, necessitating different dose regimens for different age groups. Levocetirizine, the active enantiomer of cetirizine, is used in the treatment of allergic rhinitis and chronic urticaria in Europe. Its pharmacokinetics and pharmacodynamics have not yet been studied prospectively in school-age children. This study was performed to investigate levocetirizine pharmacokinetic disposition and pharmacodynamics in relation to skin reactivity to histamine in children aged 6 to 11 years. Blood samples were obtained at predose baseline and at defined intervals up to and including 28 h after a 5-mg levocetirizine dose. Concurrently, epicutaneous tests with histamine phosphate, 1 mg/mL, were performed. Wheals and flares were traced at 10 min, and the areas were measured with a computerized digitizing system. In children aged 8.6 ± 0.4 years (± SEM), the peak levocetirizine concentration was 450 ± 37 ng/mL, and the time at which peak concns. occurred was  $1.2 \pm 0.2 h$ . terminal elimination half-life was  $5.7 \pm 0.2 \text{ h}$ , the oral clearance was 0.82  $\pm$  0.05 mL/min/kg, and the volume of distribution was  $0.4 \pm 0.02$  L/kg. Compared with predose areas, the wheals and flares produced by histamine phosphate were significantly decreased from 1 to 28 h, inclusive (P < .05). Mean maximum inhibition of wheals and flares occurred from 2 to 10 h (97%  $\pm$  1%) and from 2 to 24 h (93% ± 1%), resp. Levocetirizine had an onset of action within 1 h and provided significant peripheral antihistaminic activity for 28 h after a single dose. Once-daily dosing may be optimal in children aged 6 to 11 years, as it is in adults.

IT 130018-77-8, Levocetirizine

RL: ADV (Adverse effect, including toxicity); PAC (Pharmacological activity); PKT (Pharmacokinetics); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(levocetirizine had significant peripheral antihistaminic activity, decreased histamine phosphate induced wheals, flares and 5mg once daily dosing may be optimal in mild allergic rhinitis children aged 6 to 11 years as in adults)

REFERENCE COUNT:

THERE ARE 31 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 5 OF 19 CAPLUS COPYRIGHT 2006 ACS on STN

31

ED Entered STN: 24 Jun 2005

ACCESSION NUMBER: 2005:548316 CAPLUS

DOCUMENT NUMBER: 143:278031

TITLE: Pharmacological management of allergic rhinitis in

the elderly: Safety issues with oral

antihistamines

AUTHOR(S): Hansen, Juga; Klimek, Ludger; Hoermann, Karl CORPORATE SOURCE: Ear, Nose and Throat Department, Mannheim

University Hospital, Mannheim, Germany

SOURCE: Drugs & Aging (2005), 22(4), 289-296

CODEN: DRAGE6; ISSN: 1170-229X

PUBLISHER: Adis International Ltd.
DOCUMENT TYPE: Journal; General Review

LANGUAGE: English

AB A review. An increasing number of elderly persons in our society experience allergic rhinoconjunctivitis. Different agents are used in the pharmacol. treatment of allergic rhinitis, with histamine H1 receptor antagonists (antihistamines) being the most

frequently prescribed class. However, drug therapy of aged persons differs to a degree from that in other age groups primarily because of quant. pharmacotherapeutic problems. The main problems are co-morbidities and polymedication, which may lead to drug-drug interactions. H1 receptor antagonists block the action of histamine at specific receptors and are available for both topical and systemic administration. First-generation H1 receptor antagonists are lipophilic and therefore may cross the blood-brain barrier; they also lack specificity for the H1 receptor. Second-generation H1 receptor antagonists have reduced capacity to cross the blood-brain barrier and greater specificity for the H1 receptor. Use of first-generation H1 receptor antagonists in the elderly should be considered carefully because of the large number of adverse effects and potential for interactions with these agents. Second-generation H1 receptor antagonists such as desloratadine, levocetirizine and ebastine provide good selective H1 receptor blockade without anticholinergic or  $\alpha$ -adrenoceptor antagonist activity. Furthermore, they inhibit proinflammatory cytokines and are safe. Second-generation H1. Receptor antagonists also offer therapeutic possibilities in patients with severe liver and/or renal dysfunction.

# IT 130018-77-8, Levocetirizine

RL: ADV (Adverse effect, including toxicity); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (second-generation H1 receptor antagonist like

levocetirizine show selective H1 receptor blockade without anticholinergic, α-adrenoceptor antagonist activity and are safe for treatment of allergic rhinitis elderly patient)

REFERENCE COUNT:

21 THERE ARE 21 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 6 OF 19 CAPLUS COPYRIGHT 2006 ACS on STN

ED Entered STN: 13 May 2005

ACCESSION NUMBER: 2005:409318 CAPLUS

DOCUMENT NUMBER: 142:451842

TITLE: Pharmaceutical product comprising a  $\beta$ -2

adrenergic agonist and an H1-receptor antagonist

INVENTOR(S): Lulla, Amar; Malhotra, Geena

PATENT ASSIGNEE(S): Cipla Limited, India; Wain, Christopher Paul

SOURCE: PCT Int. Appl., 25 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT N	10.			KIN	D :	DATE		i	APPL:	ICAT:	ION 1			DA	ATE
WO 2005		A1	A1 20050512 AM, AT, AU, AZ,				йO 2								
W:	ΑE,	AG,	AL,	AM,	ΑT,	ΑU,	ΑZ,	BA,	BB,	ΒG,	BR,	BW,	BY,	ΒZ,	CA,
	CH,	CN,	co,	CR,	CU,	CZ,	DE,	DK,	DM,	DΖ,	EC,	EE,	EG,	ES,	FI,
	GB,	GD,	GE,	GH,	GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	ΚE,	KG,	KP,
	KR,	KZ,	LC,	LK,	LR,	LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,
	MX,	MZ,	NA,	NI,	NO,	NZ,	OM,	PG,	PH,	PL,	PT,	RO,	RU,	SC,	SD,
	SE,	SG,	SK,	SL,	SY,	ТJ,	TM,	TN,	TR,	TT,	ΤZ,	UA,	ŪG,	US,	UZ,
	VC,	VN,	YU,	ZA,	ZM,	ZW									
RW:	BW,	GH,	GM,	KE,	LS,	MW,	ΜZ,	NA,	SD,	SL,	SZ,	TZ,	UG,	ZM,	ZW,
	AM,	AZ,	BY,	KG,	KZ,	MD,	RU,	ТJ,	TM,	AT,	BE,	BG,	CH,	CY,	CZ,

DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

PRIORITY APPLN. INFO.:

IN 2003-MU1120 A 20031022

A pharmaceutical composition comprising at least one AR therapeutically selective isomer of a  $\beta$ -2-adrenergic agonist, or a salt, solvate, physiol. functional derivative or prodrug thereof, and at least one therapeutically selective isomer of an H1-receptor antagonist, or a salt, solvate, physiol. functional derivative or prodrug thereof, together with a pharmaceutically acceptable carrier or excipient is disclosed.

130018-77-8, Levocetirizine

RL: PEP (Physical, engineering or chemical process); PYP (Physical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)

(pharmaceutical product comprising a  $\beta2$  adrenergic agonist and an H1-receptor antagonist)

REFERENCE COUNT:

THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

ANSWER 7 OF 19 CAPLUS COPYRIGHT 2006 ACS on STN L9

Entered STN: 06 May 2005

2005:388517 CAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER: 143:19206

Randomized, double-blind, crossover comparison TITLE:

between two levocetirizine formulations

on histamine-induced cutaneous response in healthy

male human adult volunteers

Usharani, P.; Naidu, M. U. R.; Reddy, K. L. N.; AUTHOR(S):

Reddy, B. P. S.; Kumar, T. Ramesh

Department of Clinical Pharmacology and CORPORATE SOURCE:

Therapeutics, Nizam's Institute of Medical

Sciences, Hyderabad, India

Journal of Applied Research (2005), 5(1), 149-159 SOURCE:

CODEN: JAROBP; ISSN: 1537-064X

PUBLISHER: Therapeutic Solutions LLC

DOCUMENT TYPE: Journal LANGUAGE: English

Cetirizine is a highly efficacious and long acting second generation AΒ H1 receptor antagonist indicated for the treatment of allergic diseases. It is a racemate mixture composed of equal amts. of S- and R-enantiomers, and the R-enantiomer, levocetirizine, carries the majority of the histamine H1-receptor-blocking activity. Recently, levocetirizine formulation has been introduced in India for the treatment of allergic rhinitis and urticaria. Objective: The aim of this study was to compare the effect of levocetirizine (Indian formulation) vs. an international brand of levocetirizine in 12 healthy male human volunteers under fasting conditions, using pharmacodynamic measure of inhibition of histamine induced wheal and flare response. Methodol.: Twelve healthy male volunteers were enrolled in this study. All volunteers gave written informed consent before entering in the study, which was approved by the Institutional Ethics Committee of Nizam's Institute of Medical Sciences. This was a balanced, randomized, double-blind, single oral dose, crossover study, where the subjects were randomized to receive either 5 mg levocetirizine reference or test formulation after overnight fast. A ten-day period was allowed

between the 2 treatment schedules to eliminate the carry over effect of earlier treatment. Wheal and flare were induced on the forearm of the trial subjects by injecting freshly prepared histamine (0.1 mL containing 2 µg) intradermally while the subject was lying comfortably with the arm resting on the bed. Ten minutes later, wheal and flare were visualized under a bright lamp. Histamine induced wheal and flare skin test was performed before and at 2 h, 4 h, 6 h, 8 h, 12 h, and 24 h after drug administration. Results: Ten minutes after intradermal injection, 2 µg of histamine produced significant wheal and flare cutaneous response in all subjects. Administration of reference and test formulations of levocetirizine significantly inhibited the histamine induced cutaneous response in all of the subjects. Maximum inhibition of histamine induced wheal response (Iw max %) with reference was 82.45% ± 8.8% and 77.9%  $\pm$  12.9% with test formulation. Maximum inhibition of histamine induced flare response (If max %) was 80%  $\pm$  4.4% and  $81.58 \pm 6.78$  with reference and test formulations resp. The area under the antihistaminic activity minus time profile curve for wheal was 2211 mm2/h  $\pm$  270 mm2/h and 2482  $\pm$  368 mm2/h with reference and test formulations, resp., and was found to be comparable. The least square mean ratio (%), T vs. R for peak activity, Imax minus percent (maximum inhibition of histamine induced wheat and flare response), area under the activity time curve (AUC0-24 mm2/h and AUC0-24 %/h) both for untransformed and log transformed data were found to be within 80% to 125% of 90% CI limits and both formulations were well tolerated. Conclusion: It can thus be concluded that the test formulation of levocetirizine tablet is bioequivalent to reference levocetirizine tablet and both formulations are equally effective and well tolerated.

ΙT 130018-77-8, Levocetirizine

RL: ADV (Adverse effect, including toxicity); PKT (Pharmacokinetics);

THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(H1 receptor antagonist levocetirizine is equally

effective and bioequivalent to reference levocetirizine on histamine-induced cutaneous response and was well tolerated in healthy human)

THERE ARE 10 CITED REFERENCES AVAILABLE FOR REFERENCE COUNT: 10

THIS RECORD. ALL CITATIONS AVAILABLE IN THE

RE FORMAT

L9 ANSWER 8 OF 19 CAPLUS COPYRIGHT 2006 ACS on STN

Entered STN: 31 Jan 2005

2005:82181 CAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER: 143:52868

Retrospective population pharmacokinetics of TITLE:

> levocetirizine in atopic children receiving cetirizine: The ETAC study

AUTHOR(S): Hussein, Ziad; Pitsiu, Maria; Majid, Oneeb;

Aarons, Leon; de Longueville, Marc; Stockis, Armel

The ETAC Study Group, Medeval Ltd, Manchester, UK CORPORATE SOURCE:

British Journal of Clinical Pharmacology (2005), SOURCE:

59(1), 28-37

CODEN: BCPHBM; ISSN: 0306-5251

PUBLISHER: Blackwell Publishing Ltd.

DOCUMENT TYPE: Journal LANGUAGE: English

To evaluate the population pharmacokinetics of levocetirizine

in young children receiving long-term treatment with

cetirizine. Data were available from a randomized, double-blind,

parallel group and placebo-controlled study of cetirizine in 343 young children between 12 and 24 mo of age at entry, who were at high risk of developing asthma, but were not yet affected (ETAC study). Infants received oral drops of cetirizine at 0.25 mg kg-1 twice daily for 18 mo. Plasma concentration of the active enantiomer levocetirizine was determined in blood samples collected at months 3, 12 and 18 (1-3 samples per child). A one-compartment open model was fitted to the data using nonlinear mixed effects modeling (NONMEM). The influence of weight, age, gender, BSA and other covariates on CL/F and V/F was evaluated. CL/F increased linearly with weight by 0.044 l h-1 kg-1 over an intercept of 0.244 l h-1, and V/F increased linearly with weight by 0.639 l kg-1. Population ests. in children with wts. of 8 and 20 kg were 0.60 and 1.13 1 h-1 for CL/F, and 5.1 and 12.8 l for V/F, resp., with interpatient variabilities of 24.4% and 14.7%. Weight-normalized ests. of CL/F and V/F were higher than in adults. The estimated relative bioavailability was 0.28 in 12% of instances of suspected noncompliance. Levocetirizine pharmacokinetics were not influenced by severe allergy or aeroallergen sensitization. Results on the effects of concomitant medications or diseases were inconclusive due to limited pos. cases. AUC55, calculated in compliant subjects using posterior ests. of the final model, was 1952 (1227-3319)  $\mu$ g l-1 h (mean, min-max), a value similar to that in adults after intake of 5 mg oral solution 2036 (1414-2827) μg 1-1 h. The model suggests that administration of levocetirizine 0.125 mg kg-1 twice daily in children 12-48 mo of age or weighing 8-20 kg yields the same exposure as in adults taking the recommended dose of 5 mg once daily.

ΙT 130018-77-8, Levocetirizine

> RL: PKT (Pharmacokinetics); BIOL (Biological study) (population pharmacokinetic model showed 0.125 mg/kg levocetirizine given twice daily in atopic 12-48 mo aged children weighing 8-20 kg with cetirizine therapy yielded

same exposure as in adult receiving 5 mg once daily)

THERE ARE 22 CITED REFERENCES AVAILABLE FOR REFERENCE COUNT: 22 THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

ANSWER 9 OF 19 CAPLUS COPYRIGHT 2006 ACS on STN L9

Entered STN: 13 Jan 2005

ACCESSION NUMBER: 2005:28330 CAPLUS

DOCUMENT NUMBER: 142:120516

Combined pharmaceutical product comprising a TITLE:

> β2 adrenoreceptor agonist and an antihistamine for the treatment of

respiratory diseases

Lulla, Amar; Malhotra, Geena INVENTOR(S):

PATENT ASSIGNEE(S): Cipla Limited, India

SOURCE: Brit. UK Pat. Appl., 18 pp.

CODEN: BAXXDU DOCUMENT TYPE: Patent

English LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
GB 2403655	A1	20050112	GB 2003-16360	20030711
WO 2005007145	A1	20050127	WO 2004-GB3004	20040709
W: AE, AG, AL,	AM, AT	, AU, AZ, BA	, BB, BG, BR, BW, BY,	BZ, CA,

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CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW

RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

PRIORITY APPLN. INFO::

GB 2003-16360

A 20030711
```

A pharmaceutical product comprises at least 1 \( \beta 2 \) adrenoreceptor AB agonist, and at least 1 antihistamine, as a combined preparation, for simultaneous sep. or sequential use in the treatment of respiratory diseases, e.g., asthma, an allergic respiratory disorder or a related disorder. The  $\beta 2$  adrenoreceptor agonist is preferably salmeterol, bambuterol, terbutaline or formoterol or a salt, solvate or physiol. functional derivative thereof, with bambuterol-HCl, being particularly preferred. The antihistamine is preferably loratadine, decarbethoxyloratidine, cetirizine or levocetirizine, or a salt, solvate or physiol. functional derivative thereof, with cetirizine-HCl or levocetirizine-HCl, being particularly preferred. Thus, a tablet formulation contained bambuterol-HCl 10.0, cetirizine-HCl 10.0, lactose 100.20, starch 50.00, colloidal silica 2.00, microcryst. cellulose 14.50, talc 1.80, Mg stearate 1.50, and Opadry White 6.00 mg and water qs.

IT 130018-77-8, Levocetirizine

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (combined pharmaceutical product comprising  $\beta 2$  adrenoreceptor agonist and antihistamine for **treatment** of respiratory diseases)

REFERENCE COUNT:

7 THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 10 OF 19 CAPLUS COPYRIGHT 2006 ACS on STN

ED Entered STN: 09 Jan 2005

ACCESSION NUMBER: 2005:16758 CAPLUS

DOCUMENT NUMBER: 142:422656

TITLE: Chronic urticaria: Etiology, management and

current and future treatment options

AUTHOR(S): Kozel, Martina M. A.; Sabroe, Ruth A.

CORPORATE SOURCE: Department of Dermatology, Red Cross Hospital,

Beverwijk, Neth.

SOURCE: Drugs (2004), 64(22), 2515-2536

CODEN: DRUGAY; ISSN: 0012-6667

PUBLISHER: Adis International Ltd.
DOCUMENT TYPE: Journal; General Review

LANGUAGE: English

AB A review. Chronic urticaria is a common condition that can be very disabling when severe. A cause for chronic idiopathic urticaria (CIU) is only infrequently identified. Potential causes include reactions to food and drugs, infections (rarely) and, apart from an increased incidence of thyroid disease, uncomplicated urticaria is not usually associated with underlying systemic disease or malignancy. About one-third of patients with CIU have circulating functional autoantibodies against the high affinity IgE receptor or against IgE,

although it is not known why such antibodies are produced, or how the presence of such antibodies alters the course of the disease or response to treatment. There are only a few publications relating to childhood urticaria, but it is probably similar to the adult form, except that adult urticaria is more common. The diagnosis is based on patient history and it is vital to spend time documenting this in detail. Extensive laboratory tests are not required in the vast majority of patients. Chronic urticaria resolves spontaneously in 30 - 55% of patients within 5 years, but it can persist for many years. Treatment is aimed firstly at avoiding underlying causative or exacerbating factors. Histamine H1 receptor antagonists remain the mainstay of oral treatment for all forms of urticaria. The newer low-sedating antihistamines desloratadine, fexofenadine, levocetirizine and mizolastine should be tried Sedating antihistamines have more adverse effects but are useful if symptoms are causing sleep disturbance. Low-dose doxepin is effective and especially suitable for patients with associated depression. There is controversy as to whether the addition of an histamine H2 receptor antagonist or a leukotriene antagonist is helpful. For CIU, second-line agents include ciclosporin (cyclosporine) [which is effective in approx. 75% of patients], short courses of oral corticosteroids, i.v. Igs and plasmapheresis, although the last two were beneficial in small trials only. Treatments for CIU with only limited or anecdotal supportive evidence include sulfasalazine, methotrexate, stanazol, rofecoxib and cyclophosphamide. The efficacy of photo (chemo) therapy is controversial. Phys. urticarias may respond to H1 receptor antagonists, although in delayed pressure urticaria, and cold, solar and aquagenic urticaria, the response may be disappointing. Second-line agents for phys. urticarias vary depending on the urticaria and most have limited supportive evidence. The potential for spontaneous resolution, the variation in the disease activity and the unpredictable nature of the disease makes the efficacy of treatments difficult to

## IT 130018-77-8, Levocetirizine

RL: PAC (Pharmacological activity); PKT (Pharmacokinetics); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (etiol., management, and current and future treatment of

patients with chronic urticaria)

REFERENCE COUNT: 231 THERE ARE 231 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 11 OF 19 CAPLUS COPYRIGHT 2006 ACS on STN

ED Entered STN: 24 Sep 2004

ACCESSION NUMBER: 2004:781562 CAPLUS

DOCUMENT NUMBER: 141:270864

TITLE: Comparison of pharmacokinetics and metabolism of

desloratadine, fexofenadine,

levocetirizine and mizolastine in humans

AUTHOR(S): CORPORATE SOURCE: Molimard, M.; Diquet, B.; Strolin Benedetti, M. Departement de Pharmacologie, Centre Hospitalier

Universitaire, Bordeaux, Fr.

SOURCE: Fundamental & Clinical Pharmacology (2004), 18(4),

399-411

CODEN: FCPHEZ; ISSN: 0767-3981

PUBLISHER: Blackwell Publishing Ltd.
DOCUMENT TYPE: Journal; General Review

LANGUAGE: English

A review. Absorption, distribution, metabolism and excretion of AB desloratadine, fexofenadine, levocetirizine, and mizolastine in humans have been compared. The time required to reach peak plasma levels (tmax) is shortest for levocetirizine (0.9 h) and longest for desloratadine (≥3 h). Steady-state plasma levels are attained after about 6 days for desloratadine, 3 days for fexofenadine, 2-3 days for mizolastine and by the second day for levocetirizine. The apparent volume of distribution is limited for levocetirizine (0.4 L/kg) and mizolastine (1-1.2 L/kg), larger for fexofenadine (5.4-5.8 L/kg) and particularly large for desloratadine (≈ 49 1/kg). Fexofenadine and levocetirizine appear to be very poorly metabolized (≈ 5 and 14% of the total **oral** dose, resp.). Desloratadine and mizolastine are extensively metabolized. After administration of 14Clevocetirizine to healthy volunteers. 85 And 13% of the radioactivity are recovered in urine and feces, resp. In contrast, feces are the preferential route of excretion for 14C-fexofenadine (80% vs. 11% of the radioactive dose in urine). The corresponding values are 41% (urine) and 47% (feces) for 14C-desloratadine, 84-95% (feces) and 8-15% (urine) for 14C-mizolastine. The absolute bioavailability is 50-65% for mizolastine: it is high for levocetirizine as the percentage of the drug eliminated unchanged in the 48 h urine is 77% of the oral dose; the estimation for fexofenadine is at least 33%: no estimation was found for desloratadine. Fexofenadine is a P-qlycoprotein (P-qp) substrate and P-qp is certainly involved both in the poor brain penetration by the compound and, at least partially, in a number of observed drug interactions. An interaction of desloratadine with P-gp has been suggested in mice, whereas the information on mizolastine is very poor. The fact that levocetirizine is a substrate of P-gp, although weak in an in vitro model, could contribute to prevent drug penetration into the brain, whereas it is unlikely to be of any clin. relevance for P-qp-mediated drug interactions. IT 130018-77-8, Levocetirizine RL: PKT (Pharmacokinetics); BIOL (Biological study) (comparison of pharmacokinetics and metabolism of desloratadine, fexofenadine, levocetirizine and mizolastine in humans) REFERENCE COUNT: THERE ARE 104 CITED REFERENCES AVAILABLE FOR 104 THIS RECORD. ALL CITATIONS AVAILABLE IN THE

RE FORMAT

ANSWER 12 OF 19 CAPLUS COPYRIGHT 2006 ACS on STN L9

Entered STN: 02 Jan 2004

2004:2869 CAPLUS ACCESSION NUMBER:

140:47583 DOCUMENT NUMBER:

Amorphous levocetririzine dihydrochloride TITLE:

compositions for treatment of allergies

INVENTOR(S):

Reddy, Manne Satyanarayana; Rajan, Srinavasan

Thirumalai; Rao, Uppala Venkata Bhaskara; Ramayya,

Vaddadi Pattabhi

PATENT ASSIGNEE(S): Reddy's Laboratories Limited, India; Reddy's

Laboratories, Inc.

SOURCE: PCT Int. Appl., 29 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

English LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

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APPLICATION NO.
     PATENT NO.
                        KIND
                                 DATE
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                                             ______
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                         A1 20031231 WO 2003-US19777
                                                                    20030623
     WO 2004000823
         W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH,
             CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD,
             GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ,
             LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ,
             NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK,
             SL, TJ, TM, TN, TR, TT, T2, UA, UG, US, UZ, VC, VN, YU, ZA,
             ZM, ZW
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR,
             NE, SN, TD, TG
     CA 2489991
                         AA
                                 20031231
                                            CA 2003-2489991
                                                                      20030623
                                20040106
                         A1
     AU 2003277855
                                             AU 2003-277855
                                                                     20030623
     US 2004132743
                         A1 20040708 US 2003-601844
                                                                     20030623
                                             CN 2003-814416
                                 20050831
                                                                     20030623
     CN 1662515
                         Α
                                             IN 2002-MA472
                                                               A 20020621
PRIORITY APPLN. INFO.:
                                                                  W 20030623
                                             WO 2003-US19777
     A process for the preparation of the amorphous form of
AB
     levocetirizine dihydrochloride is described. A pharmaceutical
     composition comprising a prophylactically or
     therapeutically effective amount of an amorphous form of
     levocetirizine dihydrochloride and pharmaceutical
     excipients is provided. The amorphous form of
     levocetirizine dihydrochloride is suitable for pharmaceutical
     purposes in the treatment of allergies, including ailments
     such as chronic and acute allergic rhinitis, allergic conjunctivitis,
     pruritus, urticaria and the like.
     130018-87-0P, Levocetirizine dihydrochloride
     RL: PEP (Physical, engineering or chemical process); PRP (Properties);
     PYP (Physical process); SPN (Synthetic preparation); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); PROC (Process);
     USES (Uses)
        (preparation of amorphous levocetririzine-2HCl for tablets and
        treatment of allergies)
IT
     130018-77-8, Levocetirizine
     RL: RCT (Reactant); RACT (Reactant or reagent)
        (preparation of amorphous levocetririzine-2HCl for tablets and
        treatment of allergies)
REFERENCE COUNT:
                          3
                                THERE ARE 3 CITED REFERENCES AVAILABLE FOR
                                THIS RECORD. ALL CITATIONS AVAILABLE IN THE
                                RE FORMAT
L9
     ANSWER 13 OF 19 CAPLUS COPYRIGHT 2006 ACS on STN
     Entered STN: 14 Sep 2003
                          2003:719308 CAPLUS
ACCESSION NUMBER:
DOCUMENT NUMBER:
                          139:240373
                          Pharmaceutical composition of a phosphodiesterase
TITLE:
                          4 (PDE4) inhibitor or a PDE3/4 inhibitor and a
                          histamine receptor antagonist for the
                          treatment of respiratory diseases
INVENTOR(S):
                          Beume, Rolf; Bundschuh, Daniela; Weimar,
                          Christian; Wollin, Stefan-lutz
```

Searcher : Shears 571-272-2528

Altana Pharma Ag, Germany

PATENT ASSIGNEE(S):

SOURCE: PCT Int. Appl., 87 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent English

LANGUAGE:

. 1

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

	PATENT NO.						DATE						ION I			DATE		
					A1		2003	0912								2	0030225	
	W:	ΑE,	AL,	ΑU,	BA,	BR,	CA,	CN,	co,	CU	J, ]	DZ,	EC,	GE,	HR,	ID,	IL,	
		IN,	IS,	JΡ,	KR,	LT,	LV,	MA,	MK,	MX	(, I	NO,	NZ,	PH,	PL,	SG,	TN,	
		UA,	US,	VN,	ΥU,	ZA,	ZW											
	RW:	AM,	ΑZ,	BY,	KG,	ΚZ,	MD,	RU,	ТJ,	TM	1, 7	ΑT,	BE,	BG,	CH,	CY,	CZ,	
		DE,	DK,	EE,	ES,	FI,	FR,	GB,	GR,	HU	J, :	ΙE,	IT,	LU,	MC,	NL,	PT,	
		SE,	SI,	SK,	TR													
CA	2478	612			AA		2003	0912		CA	20	03-2	24780	612		2	0030225	
AU	2003	2122	68		A1		2003	0916		ΑU	20	03-2	21220	68		2	0030225	
EP	1482	938			A1		2004	1208		ΕP	200	03-	70813	30		2	0030225	
	R:	AT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GF	₹, :	IT,	LI,	LU,	NL,	SE,	MC,	
		PT,	IE,	SI,	LT,	LV,	FI,	MK,	CY,	ΑI	i, :	TR,	ВG,	CZ,	EE,	ΗU,	SK	
BR	2003	0082	20		A		2005	0104		BR	20	03-8	8220			2	0030225	
US	2005	1120	69		A1		2005	0526		US	200	03-5	50681	75		2	0030225	
JP	2005	5246	66		Т2		2005	0818		JΡ	20	03-5	5725	72		2	0030225	
NO	2004	0042	30		Α		2004	1206		NO	20	04-4	4230			2	0041006	
PRIORITY	Y APP	LN.	INFO	.:						ΕP	20	02-4	4987			A 2	0020306	
									,	wo	20	03-1	EP18	76	1	w 2	0030225	

- AB The invention discloses the combined administration of PDE4 or PDE3/4 inhibitors and histamine receptor antagonists for the treatment of respiratory diseases.
- IT 130018-77-8 130018-77-8D, Levocetirizine,
   derivs.

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(phosphodiesterase 4 (PDE4) inhibitor or PDE3/4 inhibitor combination with histamine receptor antagonist for

treatment of respiratory disease)

6

REFERENCE COUNT:

THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 14 OF 19 CAPLUS COPYRIGHT 2006 ACS on STN

ED Entered STN: 12 Feb 2003

ACCESSION NUMBER: 2003:108697 CAPLUS

DOCUMENT NUMBER:

138:162870

TITLE:

Comparative pharmacology of H1 antihistamines:

clinical relevance

AUTHOR(S):

Simons, F. Estelle R.

CORPORATE SOURCE: Section of Allergy and Clinical Immunology,

Department of Pediatrics and Child Health, Faculty of Medicine, University of Manitoba, Winnipeg, MB,

Can.

SOURCE:

American Journal of Medicine (2002), 113(9A),

38S-46S

CODEN: AJMEAZ; ISSN: 0002-9343

PUBLISHER: DOCUMENT TYPE: Excerpta Medica, Inc.
Journal; General Review

LANGUAGE: English

A review. H1 antihistamines have similar efficacy in the treatment of allergic disorders; however, they differ in terms of their chemical structure, clin. pharmacol., and safety. This review focuses on the clin. pharmacol. (pharmacokinetics and pharmacodynamics) of the newer oral H1 antihistamines (acrivastine, cetirizine, desloratadine, ebastine, fexofenadine, levocetirizine, loratadine, and mizolastine). Understanding the pharmacokinetics and pharmacodynamics of these H1 antihistamines provides an objective basis for selection of appropriate dosages and dose intervals. Pharmacokinetic and pharmacodynamic studies provide a rationale for the modified dosage regimens that may be required in special populations, such as the very young, the elderly, those with hepatic or renal dysfunction, or those taking other medications concurrently. Many H1 antihistamines are currently available for use. Clin. pharmacol. studies help physicians to select the best H1 antihistamines for their patients.

REFERENCE COUNT:

THERE ARE 28 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

ANSWER 15 OF 19 CAPLUS COPYRIGHT 2006 ACS on STN Ь9

28

Entered STN: 10 Jan 2003

ACCESSION NUMBER: 2003:22666 CAPLUS

DOCUMENT NUMBER: 138:61370

Tablets comprising cetirizine and TITLE:

pseudoephedrine

UCB, S.A., Belg.

INVENTOR(S): Fanara, Domenico; Guichaux, Anthony; Berwaer,

Monique; Deleers, Michel

PATENT ASSIGNEE(S):

PCT Int. Appl., 25 pp. SOURCE:

CODEN: PIXXD2

DOCUMENT TYPE:

English

Patent

LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PAT	PATENT NO.				KIND DATE			APPLICATION NO.						DATE		
WO	2003	0020	98		A1	_	2003	0109	1	WO 2	002-	EP63	42		2	0020610
	W:	ΑE,	AG,	AL,	AM,	AT,	ΑU,	ΑZ,	BA,	BB,	BG,	BR,	BY,	ΒZ,	CA,	CH,
		CN,	co,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	ΕE,	ES,	FI,	GB,	GD,
		GE,	GH,	GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	ΚE,	KG,	ΚP,	KR,	KZ,
		LC,	LK,	LR,	LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,
		NO,	ΝZ,	OM,	PH,	PL,	PT,	RO,	RU,	SD,	SE,	SG,	SI,	SK,	SL,	TJ,
		TM,	TN,	TR,	TT,	ΤZ,	UA,	UG,	US,	UΖ,	VN,	YU,	ZA,	ZM,	zw	
	RW:	GH,	GM,	KE,	LS,	MW,	MZ,	SD,	SL,	SZ,	TZ,	UG,	ZM,	ZW,	ΑT,	BE,
		CH,	CY,	DE,	DK,	ES,	FΙ,	FR,	GB,	GR,	ΙE,	IT,	LU,	MC,	NL,	PT,
		SE,	TR,	BF,	ВJ,	CF,	CG,	CI,	CM,	GΑ,	GN,	GQ,	GW,	ML,	MR,	NE,
		SN,	TD,	TG												
CA	2451	519			AA		2003	0109	1	CA 2	002-	2451	519		2	0020610
	2004															0020610
EΡ	1404	304			A1		2004	0407		EP 2	002-	7431	73		2	0020610
	R:	AT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	IT,	LI,	LU,	NL,	SE,	MC,
		PT,			•		FI,	-	•	•	•					
	1520														2	0020610
BR	2002	0106	50		Α		2004	1005		BR 2	002-	1065	0		2	0020610
ΝZ	5302	89			Α		2004	1126		NZ 2	002-	5302	89		2	0020610
JP	2004	5368	29		Т2		2004	1209		JP 2	003-	5083	37		2	0020610

ZA 2003009720 BG 108452	A A	20041215 20050228		2003-9720 2003-108452		20031215
US 2004170690 US 7014867	A1 B2	20040902	US	2003-481264		20031219
NO 2003005798 US 2006034928 PRIORITY APPLN. INFO.:	A A1	20040227 20060216	US	2003-5798 2005-251895 2001-115807	A	20031223 20051018 20010628
			US	2001-301250P	P	20010628
			WO	2002-EP6342	W	20020610
			US	2003-481264	А3	20031219

AB The present invention concerns a tablet comprising 2 distinct segments. More particularly the invention relates to combinations of 2 pharmaceutical substances and methods of treatment of allergic disorders. A phase 1. opened, randomized pilot study compared the oral bioavailability of exptl. 120 mg sustained-release segment pseudoephedrine formulations comprised pseudoephedrine-HCl 120, HPMC 200, microcryst. cellulose 74, colloidal SiO2 2, and Mg stearate 4 mg/tablet. This formulation was bioequivalent to the reference formulation.

IT 130018-77-8, LevoCetirizine

RL: PKT (Pharmacokinetics); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(tablets comprising cetirizine and pseudoephedrine)

REFERENCE COUNT:

THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 16 OF 19 CAPLUS COPYRIGHT 2006 ACS on STN

ED Entered STN: 16 Apr 2001

ACCESSION NUMBER: 2001:267244 CAPLUS

DOCUMENT NUMBER: 135:205289

TITLE: Effect of cetirizine, levocetirizine,

and dextrocetirizine on histamine-induced nasal

response in healthy adult volunteers

AUTHOR(S): Wang, D. Y.; Hanotte, F.; De Vos, C.; Clement, P.

CORPORATE SOURCE: National University of Singapore, Singapore,

Singapore

SOURCE: Allergy (Copenhagen) (2001), 56(4), 339-343

CODEN: LLRGDY; ISSN: 0105-4538

PUBLISHER: Munksgaard International Publishers Ltd.

DOCUMENT TYPE: Journal LANGUAGE: English

AB A randomized, double-blind, 4-way, crossover study was conducted to assess the effect of treatment with 5 mg

levocetirizine, 5 mg dextrocetirizine, 10 mg cetirizine and placebo on histamine-induced changes in the nasal airways of healthy volunteers. Four hours after a single oral intake of the drugs, the subjects were challenged by nasal aerosol application of histamine with increasing doubling concns. (from 0.25 to 32 mg/mL). Nasal resistance was measured by passive anterior rhinomanometry, and

changes in histamine threshold were calculated, together with the absolute number

of sneezes after each challenge. Both levocetirizine and cetirizine attenuated the histamine-induced increase in nasal airway resistance by nearly 50% and they concomitantly increased the

histamine threshold by 4-fold (from 8 to 32 mg/mL), compared with placebo. Sneezing was also attenuated by both levocetirizine and cetirizine. However, these antihistaminic effects were not produced by dextrocetirizine. This study shows a similar activity of levocetirizine and cetirizine in inhibiting the histamine-induced increase in nasal resistance, indicating that the antihistaminic properties of cetirizine are probably attributable to levocetirizine.

IT 130018-77-8, Levocetirizine

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(cetirizine, levocetirizine, and dextrocetirizine effect on histamine-induced nasal resistance in humans)

REFERENCE COUNT:

THERE ARE 18 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 17 OF 19 CAPLUS COPYRIGHT 2006 ACS on STN

18

ED Entered STN: 07 Feb 2001

ACCESSION NUMBER: 2001:87484 CAPLUS

DOCUMENT NUMBER:

135:116792

TITLE:

A randomized, double-blind, crossover comparison

among [the effects of] cetirizine, levocetirizine, and ucb 28557 on

histamine-induced cutaneous responses in healthy

adult volunteers

AUTHOR(S):

SOURCE:

Devalia, J. L.; De Vos, C.; Hanotte, F.; Baltes,

Ε.

CORPORATE SOURCE:

Academic Respiratory Medicine, St Bartholomew's and the Royal London School of Medicine and Dentistry, St Bartholomew's Hospital, London, UK

Allergy (Copenhagen) (2001), 56(1), 50-57

Affergy (Copenhagen) (2001), 30(1), 30-

CODEN: LLRGDY; ISSN: 0105-4538

PUBLISHER: Munksgaard International Publishers Ltd.

DOCUMENT TYPE: Journal LANGUAGE: English

The pharmacol. activity and potency of the two enantiomers of AΒ cetirizine in the management of allergic skin conditions were investigated by studying the effect of treatment with 5.0 mg cetirizine, 2.5 mg levocetirizine (the (R)-enantiomer), and 2.5 mg ucb 28557 (the (S)-enantiomer) on histamine-induced wheal and flare response in healthy volunteers. Each treatment was administered as a single oral dose in randomized, double-blind, and crossover manner, and the efficacy of treatment was assessed for 32 h as percent inhibition of the histamine-induced wheal and flare areas before treatment. Blood and urine samples were collected and analyzed for the total amts. of each drug, to elucidate their pharmacokinetic profiles. Both cetirizine and levocetirizine caused a marked inhibition of histamine-induced wheal and flare, whereas ucb 28557 was inactive. Inhibition of the wheal response by cetirizine and levocetirizine was apparent by 1 h after treatment and lasted for mean durations of 24.4 and 28.4 h, resp. In addition, the response to cetirizine and levocetirizine became maximal by 6 h after treatment, rising to 79.5% and 83.8%, resp. Similarly, cetirizine and levocetirizine also markedly inhibited the histamine-induced flare response. This effect was evident for both drugs by 1 h after administration and lasted for a

mean period of 28.4 and 26.0 h for cetirizine and levocetirizine, resp. The inhibitory effect of these compds. on histamine-induced flare response was also maximal by approx. 6 h after treatment, peaking at 88.5% and 83.6%, resp. Statistical evaluation showed that cetirizine and levocetirizine were equivalent for maximum inhibition of histamine-induced wheal and flare. However, levocetirizine was superior to cetirizine when the areas under the curve were compared. In contrast, ucb 28557 did not inhibit histamine-induced wheal and flare responses at any time during the study. Plasma concns. of levocetirizine were approx. double those of ucb 28557 at 4 and 8 h after administration, and 50-60% of the drugs were excreted unchanged in urine within 32 h. The finding that, in this model, 2.5 mg levocetirizine has comparable antihistaminic activity to 5 mg cetirizine, whereas its other enantiomer, ucb 28557, has no pharmacodynamic effect, suggests that the antihistaminic properties of cetirizine observed in the management of allergic skin conditions are likely to be attributable to levocetirizine.

#### TΨ 130018-77-8, Levocetirizine

RL: BAC (Biological activity or effector, except adverse); BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)

(cetirizine, levocetirizine, and ucb 28557 effects on histamine-induced cutaneous responses in humans)

REFERENCE COUNT:

THERE ARE 25 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 18 OF 19 CAPLUS COPYRIGHT 2006 ACS on STN

25

Entered STN: 02 Feb 1998 F.D

ACCESSION NUMBER: 1998:62646 CAPLUS

DOCUMENT NUMBER: 128:80008

TITLE:

Pharmaceutical compositions for the

treatment of rhinitis containing

diphenylmethylpiperazinylacetic acid or amide

derivatives

Van de Venne, Herman; Martin, Jean-Pierre INVENTOR(S):

PATENT ASSIGNEE(S): Ucb, S.A., Belg.

SOURCE:

Brit. UK Pat. Appl., 17 pp.

CODEN: BAXXDU

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
GB 2311940	A1	19971015	GB 1997-6842	19970404
GB 2311940	В2	20000719		
US 6469009	В1	20021022	US 1996-629144	19960408
AU 9717740	A1	19971016	AU 1997-17740	19970404
AU 723364	В2	20000824		
BR 9701686	Α	19981110	BR 1997-1686	19970407
US 2001020023	A1	20010906	US 2001-838190	20010420
US 6489329	В2	20021203		
PRIORITY APPLN. INFO.:			US 1996-629144	A <sub>2</sub> 19960408

OTHER SOURCE(S):

MARPAT 128:80008

A pharmaceutical composition comprising a therapeutically

effective amount of a mixture consisting essentially of (1) a compound selected from pseudoephedrine, phenylpropanolamine and phenylephrine, an individual optical isomer or a pharmaceutically acceptable salt thereof, and (2) at least one compound selected from 2-[4(diphenylmethyl)-1-piperazinyl]-acetic acid or amide derivs., an individual optical isomer or a pharmaceutically acceptable salt thereof. A capsule contained cetirizine 5, and pseudoephedrine, 120 mg. Patients suffering from allergic rhinitis were administered one capsule in the morning and one in the evening for 3 wk. After 3 wk administration of the capsules no more symptoms were observed in 53% of the patients.

ΙT 130018-77-8

> RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(pharmaceutical compns. for treatment of rhinitis containing diphenylmethylpiperazinylacetic acid or amide derivs.)

ANSWER 19 OF 19 CAPLUS COPYRIGHT 2006 ACS on STN L9

Entered STN: 11 Jun 1994 ED

ACCESSION NUMBER: 1994:307492 CAPLUS

DOCUMENT NUMBER: 120:307492

TITLE: Pharmaceutical compositions containing optically

pure (+) cetirizine for the treatment of

allergic disorders

INVENTOR(S): Gray, Nancy M.

PATENT ASSIGNEE(S): Sepracor, Inc., USA PCT Int. Appl., 36 pp. SOURCE:

CODEN: PIXXD2

DOCUMENT TYPE: Patent English LANGUAGE:

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

	TENT											TION			D.	ATE
										WO	1993	-US89	99		1	9930922
												, DK,				
												, NL,				
		RO,	RU,	SD,	SE,	SK,	UA									
	RW:	AT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR	, IE	, IT,	LU,	MC,	NL,	PT,
												, MR,				
AU	9351	361			A1		1994	0412		AU	1993	-5136	1		1	9930922
										EΡ	1994	-9102	60		1	9930922
EP	6619	75			В1		1999	0317								
	R:	ΑT,	BE,	CH,	DĒ,	DK,	ES,	FR,	GB,	GR	, IE	, IT,	LI,	LU,	MC,	NL,
			SE													
JP	0850	1562			Т2		1996	0220		JΡ	1993	-5084	28		1	9930922
EP	8856	11			A2		1998	1223		ΕP	1998	-1155	19		1	9930922
EP	8856	11			A3		1999	0107								
	R:	AT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR	, IT	, LI,	LU,	NL,	SE,	MC,
		PT,														
ΑT	1776	36			E							-9102				9930922
	2128											-9102				9930922
US	5627											-6226				9960326
	9859						1998	0521		ΑU	1998	-5934	2		1	9980317
AU	7036	90					_	0401								
RIORIT	Y APP	LN.	INFO	.:						US	1992	-9509	10		A 1	9920924

EP 1994-910260 A3 19930922 WO 1993-US8999 W 19930922 US 1993-167722 B1 19931215 Pharmaceutical compns. containing optically pure (+) cetirizine (I) are AB used for the treatment of seasonal and perennial allergic rhinitis in humans while avoiding the concomitant liability of adverse effects associated with the racemic mixture of cetirizine. The optically pure (+) isomer is also useful for the treatment of allergic asthma and chronic and phys. urticaria. A capsule contained I 2.0, lactose 103.75, cornstarch 18.75, and Mg stearate 0.05 mg. 130018-77-8 130018-87-0 IT RL: BIOL (Biological study) (pharmaceutical compns. containing, for treatment of allergic disorders) FILE 'MEDLINE' ENTERED AT 16:16:58 ON 10 APR 2006 FILE 'BIOSIS' ENTERED AT 16:16:58 ON 10 APR 2006 Copyright (c) 2006 The Thomson Corporation FILE 'EMBASE' ENTERED AT 16:16:58 ON 10 APR 2006 Copyright (c) 2006 Elsevier B.V. All rights reserved. FILE 'WPIDS' ENTERED AT 16:16:58 ON 10 APR 2006 COPYRIGHT (C) 2006 THE THOMSON CORPORATION FILE 'CONFSCI' ENTERED AT 16:16:58 ON 10 APR 2006 COPYRIGHT (C) 2006 Cambridge Scientific Abstracts (CSA) FILE 'SCISEARCH' ENTERED AT 16:16:58 ON 10 APR 2006 Copyright (c) 2006 The Thomson Corporation FILE 'JICST-EPLUS' ENTERED AT 16:16:58 ON 10 APR 2006 COPYRIGHT (C) 2006 Japan Science and Technology Agency (JST) FILE 'JAPIO' ENTERED AT 16:16:58 ON 10 APR 2006 COPYRIGHT (C) 2006 Japanese Patent Office (JPO) - JAPIO L10 416 SEA ABB=ON PLU=ON L6 3 SEA ABB=ON PLU=ON L10 AND (EXCIPIENT OR (STABILIS? OR L11 STABILIZ? OR SUSPEND? OR SUSPENS?) (5A) AGENT) 258 SEA ABB=ON PLU=ON L10 AND (L2 OR CETIRIZINE) L12 4 SEA ABB=ON PLU=ON L12 AND (CRYSTAL? OR CRYST## OR L13 AMORPH?) 121 SEA ABB=ON PLU=ON L10 AND (ORAL? OR PER OS OR MOUTH) (S) (A L14 DMIN? OR DOSAGE OR DOSE OR DOSING OR INTAK?) 5 SEA ABB=ON PLU=ON L14 AND (TABLET OR PILL OR CAPSUL? OR L15 SOLID?) 17 SEA ABB=ON PLU=ON L10 AND (TABLET OR PILL OR CAPSUL? OR L16 SOLID?) 18 SEA ABB=ON PLU=ON L11 OR L13 OR L15 OR L16 L17 17 DUP REM L17 (1 DUPLICATE REMOVED) L18 L18 ANSWER 1 OF 17 WPIDS COPYRIGHT 2006 THE THOMSON CORP on STN

Searcher : Shears 571-272-2528

ACCESSION NUMBER: 2005-768522 [78] WPIDS

C2005-234980

DOC. NO. CPI:

TITLE:

Pharmaceutical composition for prevention of hair loss and for treatment of female

hirsutism and benign prostatic hyperplasia, comprises

as active ingredient, 2-(2-(4-((4chlorophenyl) phenylmethyl) -1-

piperazinyl) ethoxy) -acetic acid.

DERWENT CLASS:

B03 D21

INVENTOR(S):

LEE, E

PATENT ASSIGNEE(S):

(LEEE-I) LEE E

COUNTRY COUNT:

109

PATENT INFORMATION:

LA PG PATENT NO KIND DATE WEEK

WO 2005099653 A1 20051027 (200578)\* EN

RW: AT BE BG BW CH CY CZ DE DK EA EE ES FI FR GB GH GM GR HU IE IS IT KE LS LT LU MC MW MZ NA NL OA PL PT RO SD SE SI SK SL SZ TR

TZ UG ZM ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BW BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EC EE EG ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KM KP KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NA

NI NO NZ OM PG PH PL PT RO RU SC SD SE SG SK SL SM SY TJ TM TN

TR TT TZ UA UG US UZ VC VN YU ZA ZM ZW

## APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
WO 2005099653	A1	WO 2005-KR1063	20050413

PRIORITY APPLN. INFO: KR 2005-29495

20050408; KR

2004-25869

20040414

AN 2005-768522 [78] WPIDS AB

WO2005099653 A UPAB: 20051205

NOVELTY - A pharmaceutical composition for the prevention of hair loss, and having the effects of epilation, hair toning, and hair growth, and for the treatment of female hirsutism and benign prostatic hyperplasia, comprises as an active ingredient, 2-(2-(4-((4chlorophenyl) phenylmethyl) -1-piperazinyl) ethoxy) acetic acid, or its salt, hydrate, or solvate.

ACTIVITY - Anti-allergic; Anti-histamine; Anti-androgenic; Depilatory; Cytostatic.

Effects for the prevention of hair loss were investigated by administering the tablet of pharmaceutical composition to 5 healthy adult males who were older than 20 years old, three times a day (30 mg=3x10 mg). The effects were superior within 1 week in all of 5 people. The newly produced hairs had color and thickness much stronger than those of old hairs.

MECHANISM OF ACTION - None given.

USE - The pharmaceutical composition is used for the prevention of hair loss, and having the effects of epilation, hair toning, and hair growth, and for the treatment of female hirsutism and benign prostatic hyperplasia. It is used in the form of a tablet, powder, dried syrup, chewable tablet, granule, chewing tablet, capsules, soft capsule, pill, drink, and sublingual tablet (claimed). It is useful to hairs in all parts of human bodies where there are hair roots and follicles, such as the hair

> Shears Searcher : 571-272-2528

roots and follicles on the head, hair on the head, inner and outer eyelashes, mustache, hair of armpit, and pubic hair.

ADVANTAGE - The pharmaceutical composition is non-steroidal, and has no side effects, such as sexual function disorder shown in the conventional oral epilation agents and prostatic hyperplasia treatment agents. It has very superior activities; and is very easy to use compared to applicable formulations owing to simple ways of administration.

DESCRIPTION OF DRAWING(S) - The figure is a graph showing the changes in the average weight of prostate glands of castrated male SD (sic) rats one week, after the oral administration of the pharmaceutical composition to measure its anti-androgenic active effects. Dwg.1/5

L18 ANSWER 2 OF 17 WPIDS COPYRIGHT 2006 THE THOMSON CORP on STN

ACCESSION NUMBER: 2005-356059 [36]

WPIDS

DOC. NO. CPI:

C2005-110146

TITLE:

Pharmaceutical composition for the

prophylaxis or treatment of

respiratory disease comprises isomer of

beta-2-adrenergic agonist and isomer of H1-receptor

antagonist.

DERWENT CLASS:

B05

INVENTOR(S):

LULLA, A; MALHOTRA, G

PATENT ASSIGNEE(S):

(CIPL-N) CIPLA LTD; (WAIN-I) WAIN C P

COUNTRY COUNT:

108

PATENT INFORMATION:

PATENT NO	KIND	DATE	WEEK	LA	PG

WO 2005041969 A1 20050512 (200536) \* EN 25

RW: AT BE BG BW CH CY CZ DE DK EA EE ES FI FR GB GH GM GR HU IE IT KE LS LU MC MW MZ NA NL OA PL PT RO SD SE SI SK SL SZ TR TZ UG ZM ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BW BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EC EE EG ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NA NI NO NZ OM PG PH PL PT RO RU SC SD SE SG SK SL SY TJ TM TN TR

TT TZ UA UG US UZ VC VN YU ZA ZM ZW 

# APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
WO 2005041969	A1	WO 2004-GB4467	20041021
IN 2003001120	I3	IN 2003-MU1120	20031022

PRIORITY APPLN. INFO: IN 2003-MU1120 20031022

AN 2005-356059 [36] WPIDS

AΒ WO2005041969 A UPAB: 20050608

> NOVELTY - A pharmaceutical composition comprises at least one selective isomer of beta -2-adrenergic agonist (a) or its salt, solvate, functional derivative or their prodrugs and at least one selective isomer of an H1-receptor antagonist (b) or its salt, solvate, functional derivative or their prodrugs.

ACTIVITY - Respiratory-Gen.; Antiasthmatic; Antiallergic.

MECHANISM OF ACTION - beta -2-Adrenergic agonist; H1-receptor antagonist.

USE - In the manufacture of a medicament for the prophylaxis or treatment of a respiratory disease in a mammal (claimed). The disease is e.g. asthma, allergic respiratory disorders.

ADVANTAGE - The composition provides an enhanced, synergistic therapeutic effect.

Dwg.0/0

L18 ANSWER 3 OF 17 WPIDS COPYRIGHT 2006 THE THOMSON CORP on STN

ACCESSION NUMBER: 2005-295844 [30] WPIDS

DOC. NO. CPI:

C2005-091351

TITLE:

Topical composition for treating rhinitis

comprises antihistamine drug and mast cell inhibitor,

non steroidal antiinflammatory drug,

phosphodiesterase inhibitor, anti immunoglobulin E agent, heparin, topical steroid or leukotriene

blocker.

DERWENT CLASS:

B05 B07 LANE, E M

INVENTOR(S):

PATENT ASSIGNEE(S): (QTMQ-N) QTM LLC

COUNTRY COUNT:

108

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG

WO 2005030331 A1 20050407 (200530) \* EN 21

RW: AT BE BG BW CH CY CZ DE DK EA EE ES FI FR GB GH GM GR HU IE IT KE LS LU MC MW MZ NA NL OA PL PT RO SD SE SI SK SL SZ TR TZ UG ZM ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BW BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EC EE EG ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NA NI NO NZ OM PG PH PL PT RO RU SC SD SE SG SK SL SY TJ TM TN TR TT TZ UA UG US UZ VC VN YU ZA ZM ZW

#### APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
WO 2005030331	A1	WO 2004-US31380	20040927

PRIORITY APPLN. INFO: US 2003-505920P 20030926

2005-295844 [30] WPIDS AN

WO2005030331 A UPAB: 20050512 AΒ

> NOVELTY - Topical pharmaceutical composition comprises an antihistamine drug and a drug composition selected from a mast cell inhibitor, a non-steroidal anti-inflammatory drug (NSAID), a phosphodiesterase inhibitor, an anti-immunoglobulin E (IgE) agent, heparin, a topical steroid or a leukotriene blocker in an excipient.

ACTIVITY - Antiallergic; Antiinflammatory.

MECHANISM OF ACTION - Histamine inhibitor; Phosphodiesterase inhibitor; IgE inhibitor; Leukotriene blocker.

USE - For treating allergic and non-allergic rhinitis (claimed).

ADVANTAGE - The combination provides a topical composition

containing a combination of antihistamine with other drugs capable of treating allergic or non-allergic rhinitis by intervening with the allergic cascade at multiple points and also provides relief from associated symptoms such as nasal itching, rhinorrhea, nasal obstruction and loss of smell not treatable with the antihistamine alone; as compared to the prior art compositions containing a single medicament such as antihistamine or steroids. This improves the response to antihistamine and greatly improves the therapeutic effect by providing superior relief from the symptoms. The topical delivery of the composition further provides improved simplicity in dosing, improved patient compliance and significant cost savings.

Dwg.0/0

L18 ANSWER 4 OF 17 WPIDS COPYRIGHT 2006 THE THOMSON CORP on STN

ACCESSION NUMBER:

2005-076438 [09] WPIDS

DOC. NO. CPI:

C2005-026655

TITLE:

Pharmaceutical product useful for treating

respiratory disease e.g. asthma comprises beta-2

adrenoreceptor agonist and antihistamine.

DERWENT CLASS:

B05

INVENTOR(S):

LULLA, A; MALHOTRA, G

PATENT ASSIGNEE(S):

(CIPL-N) CIPLA LTD; (WAIN-I) WAIN C P

COUNTRY COUNT: 108

PATENT INFORMATION:

PAT	ENT	ИО			KI	ND I	DAT	<b>Ξ</b>	V	VEE	ζ		LA		?G							
GB	240	3655	5		Α	20	050	112	(20	0050	9),	ŧ		18								
WO	200	5001	7145	5	Α1	200	050:	127	(20	005:	10)	Eì	1									
	RW:	ΑT	ΒE	BG	BW	CH	CY	CZ	DE	DK	EΑ	EΕ	ES	FI	FR	GB	GH	GM	GR	HU	ΙE	ΙT
		KE	LS	LU	MC	MW	MZ	NA	NL	OA	PL	PT	RO	SD	SE	SI	sĸ	$\mathtt{SL}$	sz	TR	TZ	UG
		ZM	ZW																			
	W:	ΑE	ΑG	AL	AM	ΑT	ΑU	ΑZ	BA	ВВ	BG	BR	BW	BY	BZ	CA	CH	CN	CO	CR	CU	CZ
		DE	DK	DM	DZ	EC	EE	EG	ES	FI	GB	GD	GE	GH	GM	HR	HU	ΙD	IL	IN	IS	JP
		KE	KG	ΚP	KR	ΚZ	LC	LK	LR	LS	LT	LU	LV	MA	MD	MG	MK	MN	MW	ΜX	MZ	NA
		NI	NO	ΝZ	OM	PG	PH	PL	PT	RO	RU	SC	SD	SE	SG	SK	$\mathtt{SL}$	SY	TJ	TM	TN	TR

## APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE		
GB 2403655	A	GB 2003-16360	20030711		
WO 2005007145	A1	WO 2004-GB3004	20040709		

PRIORITY APPLN. INFO: GB 2003-16360 20030711

TT TZ UA UG US UZ VC VN YU ZA ZM ZW

AN 2005-076438 [09] WPIDS

AB GB 2403655 A UPAB: 20050207

NOVELTY - Pharmaceutical product comprises at least one beta -2 adrenoreceptor agonist and at least one antihistamine.

ACTIVITY - Respiratory-Gen.; Antiasthmatic; Antiallergic; Antiinflammatory.

MECHANISM OF ACTION - None given.

USE - For the prophylaxis or treatment of a respiratory disease in mammal e.g. human (claimed); for treating asthma.

ADVANTAGE - The combination of beta -2 adrenoreceptor agonist and

antihistamine alleviates allergic rhinitis and mild to moderate asthma symptoms. The composition provides enhanced synergistic effect. Dwg.0/0

L18 ANSWER 5 OF 17 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on

STN

ACCESSION NUMBER: 2005:284295 BIOSIS DOCUMENT NUMBER: PREV200510066148

TITLE: Clinical randomized controlled trials of

levocetirizine in treatment of 72 patients with anaphylactic rhinitis.

AUTHOR(S): Wang De-hui [Reprint Author]; Zheng Hai-hui; Wang

Zheng-min; Wang Jia-dong; Li Ming; Wang Jia-bin; Li

Ji-ping; Cao Yi; Ma Zhong-chao

CORPORATE SOURCE: Fudan Univ, Affiliated Ophthalmol and Otorhinolaryngol

Hosp, Shanghai 200031, Peoples R China

SOURCE: Zhongguo Xinyao yu Linchuang Zazhi, (MAY 2005) Vol. 24,

No. 5, pp. 366-369.

ISSN: 1007-7669.

DOCUMENT TYPE: Article LANGUAGE: Chinese

ENTRY DATE: Entered STN: 27 Jul 2005

Last Updated on STN: 27 Jul 2005

AB AIM: To evaluate the efficacy and safety of levocetirizine

and cetirizine in the treatment of perennial allergic

rhinitis. METHODS: One hundred and forty-four patients between 18-65

a were randomly divided into treatment group and control

group in a multi-center, double-blind, randomized active-controlled

trial. Seventy-two patients in treatment group were

treated with levocetirizine tablets (5 mg,

po, qd), 72 patients in control group were treated with

cetirizine tablets (10 mg, po, qd), all together for 14 d: RESULTS: Seventy patients of the either two groups were completed with

the trial. The total clinical effective rates were 89% and 83% for the **treatment** group after 7 d and 14 d, and those of the control group were 94% and 93%, respectively. There was no

control group were 94% and 93%, respectively. There was no statistical difference in the symptom integral lowering index (P >

0.05). No severe adverse reactions occurred both in treatment

group and control group. The incidence of adverse reaction rates 118 were for levocetinzine and 18% in cetirizine. The oscitancy rate of

treatment group was 6%, anal 11% for control group. There was no clinical diversification correlatively. CONCLUSION:

Levocetirizine is effective and safe in the treatment of perennial allergic rhinitis, nearly similar to cetirizine.

L18 ANSWER 6 OF 17 EMBASE COPYRIGHT (c) 2006 Elsevier B.V. All rights reserved on STN

ACCESSION NUMBER: 2005168928 EMBASE

TITLE: Randomized, double-blind, crossover comparison between

two levocetirizine formulations on

histamine-induced cutaneous response in healthy male

human adult volunteers.

AUTHOR: Usharani P.; Naidu M.U.R.; Reddy K.L.N.; Reddy B.P.S.;

Kumar T.R.

CORPORATE SOURCE: Dr. P. Usharani, Dept. of Clin. Pharmacol./Therapeut.,

Nizam's Inst. of Medical Sciences, Hyderabad, India

SOURCE: Journal of Applied Research, (2005) Vol. 5, No. 1, pp.

149-159. . Refs: 11

ISSN: 1537-064X CODEN: JAROBP

COUNTRY: United States
DOCUMENT TYPE: Journal; Article
FILE SEGMENT: 030 Pharmacology

037 Drug Literature Index 038 Adverse Reactions Titles

LANGUAGE: English
SUMMARY LANGUAGE: English

ENTRY DATE: Entered STN: 5 May 2005

Last Updated on STN: 5 May 2005

Cetirizine is a highly efficacious and long acting second generation AΒ H1 receptor antagonist indicated for the treatment of allergic diseases. It is a racemate mixture composed of equal amounts of S- and R-enantiomers, and the R-enantioner, levocetirizine , carries the majority of the histamine H1-receptor-blocking activity. Recently, levocetirizine formulation has been introduced in India for the treatment of allergic rhinitis and urticaria. Objective: The aim of this study was to compare the effect of levocetirizine (Indian formulation) versus an international brand of levocetirizine in 12 healthy male human volunteers under fasting conditions, using pharmacodynamic measure of inhibition of histamine induced wheal and flare response. Methodology: Twelve healthy male volunteers were enrolled in this study. All volunteers gave written informed consent before entering in the study, which was approved by the Institutional Ethics Committee of Nizam's Institute of Medical Sciences. This was a balanced, randomized, double-blind, single oral dose, crossover study, where the subjects were randomized to receive either 5 mg levocetirizine reference or test formulation after overnight fast. A ten-day period was allowed between the 2 treatment schedules to eliminate the carry over effect of earlier treatment. Wheal and flare were induced on the forearm of the trial subjects by injecting freshly prepared histamine (0.1 mL containing 2 µg) intradermally while the subject was lying comfortably with the arm resting on the bed. Ten minutes later, wheal and flare were visualized under a bright lamp. Histamine induced wheal and flare skin test was performed before and at 2 hours, 4 hours, 6 hours, 8 hours, 12 hours, and 24 hours after drug administration. Results: Ten minutes after intradermal injection, 2 µq of histamine produced significant wheal and flare cutaneous response in all subjects. Administration of reference and test formulations of levocetirizine significantly inhibited the histamine induced cutaneous response in all of the subjects. Maximum inhibition of histamine induced wheal response (I(w) max %) with reference was  $82.45\% \pm 8.8\%$  and  $77.9\% \pm 12.9\%$  with test formulation. Maximum inhibition of histamine induced flare response (I (f) max %) was 80%  $\pm$  4.4% and 81.58  $\pm$  6.7% with reference and test formulations respectively. The area under the antihistaminic activity minus time profile curve for wheal was 2211 mm(2)/hr ± 270 mm(2)/hr and 2482  $\pm$  368 mm(2)/hr with reference and test formulations, respectively, and was found to be comparable. The least square mean ratio (%),T versus R for peak activity, Imax minus percent (maximum inhibition of histamine induced wheal and flare response), area under the activity time curve (AUC(0-24) mm(2)/hr and AUC(0-24)%/hr) both for untransformed and log transformed data were found to be within 80%; to 125% of 90% CI limits and both formulations were well tolerated. Conclusion: It can thus be concluded that the test formulation of levocetirizine tablet is bioequivalent to reference levocetirizine tablet and both formulations are equally effective and well tolerated.

L18 ANSWER 7 OF 17 EMBASE COPYRIGHT (c) 2006 Elsevier B.V. All rights reserved on STN

ACCESSION NUMBER: 2005122888 EMBASE

TITLE: Levocetirizine, new indication.

AUTHOR: Mealy N.E.; Bayes M.

CORPORATE SOURCE: N.E. Mealy, Prous Science, P.O. Box 540, 08080

Barcelona, Spain

SOURCE: Drugs of the Future, (2005) Vol. 30, No. 1, pp. 86-87.

Refs: 3

ISSN: 0377-8282 CODEN: DRFUD4

COUNTRY: Spain

DOCUMENT TYPE: Journal; Note

FILE SEGMENT: 011 Otorhinolaryngology

015 Chest Diseases, Thoracic Surgery and

Tuberculosis

030 Pharmacology

037 Drug Literature Index 038 Adverse Reactions Titles

LANGUAGE:

English

ENTRY DATE: Entered STN: 7 Apr 2005

Last Updated on STN: 7 Apr 2005

DATA NOT AVAILABLE FOR THIS ACCESSION NUMBER

L18 ANSWER 8 OF 17 WPIDS COPYRIGHT 2006 THE THOMSON CORP on STN

ACCESSION NUMBER:

2004-461088 [43] WPIDS

DOC. NO. CPI:

C2004-172229

TITLE:

New crystalline and amorphous

forms of dextro/levo rotatory dihydrochloride salt of cetrizine are histamine receptor antagonists useful

in the treatment of e.g. chronic and acute allergic rhinitis, allergic conjunctivitis and

urticaria.

DERWENT CLASS:

B03

INVENTOR(S):

JOGA, R; REDDY, M S; SRINIVASAN, T R; UPPALA, V B R; VADDADI, P R; ERPARA, V B R; JORGAR, R; SREENIVASAN, T R; WADADDY, P R; MANNE, S R; PRASAD, T R; RAJENDER,

J; RAO UPPALA, V B

PATENT ASSIGNEE(S):

(REDD-N) REDDY'S LAB LTD; (REDD-N) REDDYS LAB LTD;

(REDD-N) DR REDDY'S LABORATORIES INC

COUNTRY COUNT:

107

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG

WO 2004050647 A2 20040617 (200443)\* EN 37

RW: AT BE BG BW CH CY CZ DE DK EA EE ES FI FR GB GH GM GR HU IE IT KE LS LU MC MW MZ NL OA PT RO SD SE SI SK SL SZ TR TZ UG ZM ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BW BY BZ CA CH CN CO CR CU CZ

DE DK DM DZ EC EE EG ES FI GB GD GE GH GM HR HU ID IL IN IS JP

KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NI

EN

NO NZ OM PG PH PL PT RO RU SC SD SE SG SK SL SY TJ TM TN TR TT TZ UA UG US UZ VC VN YU ZA ZM ZW

US 2004186112 A1 20040923 (200463)

AU 2003297640 Al 20040623 (200472)

IN 2002000908 I4 20050304 (200555) CN 1692105 A 20051102 (200622)

## APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
WO 2004050647	A2	WO 2003-US38494	20031204
US 2004186112	A1	US 2003-729856	20031204
AU 2003297640	A1	AU 2003-297640	20031204
IN 2002000908	I4	IN 2002-CH908	20021204
CN 1692105	Α	CN 2003-100543	20031204

#### FILING DETAILS:

PATENT NO	KIND	PATENT NO
AU 2003297640	Al Based on	WO 2004050647

PRIORITY APPLN. INFO: IN 2002-CH908 20021204

AN 2004-461088 [43] WPIDS

AB W02004050647 A UPAB: 20050902

NOVELTY - Crystalline (A) and amorphous (C) forms of dextrorotatory dihydrochloride salt of (2-(4-((4-chlorophenyl)-phenyl methyl)-1-piperazinyl) ethoxy)

acetic acid (cetirizine), crystalline (B)

and amorphous (D) forms of levorotatory dihydrochloride salt of cetirizine is new.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for:

- (1) preparation of (A), (B), (C) and (D);
- (2) a composition (I) comprising dextrorotatory dihydrochloride salt of cetirizine as a solid (where at least 80 weight% of dextrorotatory dihydrochloride salt of cetirizine is in an amorphous form); and
- (3) a composition (II) comprising levorotatory dihydrochloride salt of **cetirizine** as a **solid** (where at least 80 weight% of the levorotatory dihydrochloride salt of **cetirizine** is in an **amorphous** form).

ACTIVITY - Antiallergic; Antiinflammatory; Ophthalmological; Antipruritic; Dermatological.

MECHANISM OF ACTION - H1 histamine receptor antagonist.

USE - The crystalline and amorphous salt

forms of cetirizine dihydrochloride are effective in the treatment of allergies e.g. chronic and acute allergic rhinitis, allergic conjunctivitis, pruritus and urticaria.

ADVANTAGE - Cetirizine provides safe and effective, symptomatic relief of seasonal allergies, and includes less sedation, low anticholinergic activity and longer acting duration.

Dwg.0/6

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ACCESSION NUMBER: 2005027529 EMBASE

TITLE: [Papers in the DAZ (Deutsche Apotheker Zeitung) -

Solid knowledge and good editing].

AUFSATZE IN DER DAZ - WISSEN GUT AUFBEREITET.

AUTHOR: Caesar W.

SOURCE: Deutsche Apotheker Zeitung, (23 Dec 2004) Vol. 144, No.

52, pp. 49-58. .

ISSN: 0011-9857 CODEN: DAZEA2

COUNTRY: Germany

DOCUMENT TYPE: Journal; General Review

FILE SEGMENT:

030 Pharmacology

037

Drug Literature Index

LANGUAGE:

German

ENTRY DATE:

Entered STN: 4 Feb 2005

Last Updated on STN: 4 Feb 2005

DATA NOT AVAILABLE FOR THIS ACCESSION NUMBER

L18 ANSWER 10 OF 17 WPIDS COPYRIGHT 2006 THE THOMSON CORP on STN

ACCESSION NUMBER: 2004-169060 [16] WPIDS

DOC. NO. CPI:

C2004-066821

TITLE:

New amorphous form of

levocetirizine dihydrochloride, useful for the treatment of chronic and acute allergic rhinitis, allergic conjunctivitis, pruritus or

urticaria.

DERWENT CLASS:

B03

INVENTOR(S):

RAJAN, S T; RAMAYYA, V P; RAO, U V B; REDDY, M S; MANNE, S R; SRINIVASAN, T R; UPPALA, V B R; VADDADI,

PR 104

PATENT ASSIGNEE(S):

(REDD-N) REDDY'S LAB LTD; (REDD-N) REDDYS LAB LTD

COUNTRY COUNT:

PATENT INFORMATION:

KIND DATE WEEK LA PG PATENT NO

WO 2004000823 Al 20031231 (200416)\* EN 29

RW: AT BE BG CH CY CZ DE DK EA EE ES FI FR GB GH GM GR HU IE IT KE LS LU MC MW MZ NL OA PT RO SD SE SI SK SL SZ TR TZ UG ZM ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EC EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NI NO NZ OM PG PH PL PT RO RU SC SD SE SG SK SL TJ TM TN TR TT TZ UA UG

US UZ VC VN YU ZA ZM ZW

US 2004132743 A1 20040708 (200445) AU 2003277855 A1 20040106 (200447) IN 2002000472 I4 20050304 (200555) AU 2003277855 A8 20040106 (200562) CN 1662515 A 20050831 (200611)

APPLICATION DETAILS:

PATENT NO KIND APPLICATION DATE \_\_\_\_\_\_ WO 2003-US19777 20030623 US 2003-601844 20030623 AU 2003-277855 20030623 IN 2002-CH472 20020621 AU 2003-277855 20030623 CN 2003-814416 20030623 WO 2004000823 A1 US 2004132743 A1 AU 2003277855 A1 IN 2002000472 I4 AU 2003277855 A8 CN 1662515 A

FILING DETAILS:

KIND PATENT NO PATENT NO AU 2003277855 Al Based on AU 2003277855 A8 Based on WO 2004000823 WO 2004000823 AU 2003277855 A8 Based on

PRIORITY APPLN. INFO: IN 2002-CH472

20020621

AN 2004-169060 [16] WPIDS

AB W02004000823 A UPAB: 20050902

NOVELTY - An amorphous form of levocetirizine

dihydrochloride ((-)-(2-(4-((4-chlorophenyl)-phenyl

methyl)-1-piperazinyl)ethoxy)acetic acid

dihydrochloride) (I) is new.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for:

(1) A composition comprising (I) as a solid (where at

least 80 (preferably at least 90, especially at least 95, particularly at least 99) weight% of (I) is in an amorphous form); and

(2) Preparation of amorphous form of (I).

ACTIVITY - Antiallergic; Antiinflammatory; Ophthalmological; Antipruritic; Dermatological.

MECHANISM OF ACTION - None given.

USE - Amorphous (I) is useful in a pharmaceutical composition (claimed) for the treatment of chronic and acute allergic rhinitis, allergic conjunctivitis, pruritus or urticaria.

ADVANTAGE - The amorphous form of (I) is free of crystalline forms of cetirizine dihydrochloride. The amorphous form of (I) is obtained by a process, which is simple, eco-friendly and cost-effective; can be easily handled in pharmaceutical processing; and provides enhanced solubility. Dwg.0/3

L18 ANSWER 11 OF 17 WPIDS COPYRIGHT 2006 THE THOMSON CORP on STN

ACCESSION NUMBER:

2004-071377 [07] WPIDS

CROSS REFERENCE: DOC. NO. CPI: 2004-071376 [07] C2004-029508

TITLE:

New amorphous form of cetirizine

dihydrochloride useful in pharmaceutical formulations

for treating allergic syndromes e.g. chronic and acute allergic rhinitis.

DERWENT CLASS:

INVENTOR(S): RA

RAJAN, S T; REDDY, M S; SHANKAR, R R; VARDHAN, S V

PATENT ASSIGNEE(S): (REDD-N) REDDY'S LAB LTD

COUNTRY COUNT:

103

PATENT INFORMATION:

PATENT NO	KIND	DATE	WEEK	LA	PG

WO 2003104212 A1 20031218 (200407) \* EN 21

RW: AT BE BG CH CY CZ DE DK EA EE ES FI FR GB GH GM GR HU IE IT KE LS LU MC MW MZ NL OA PT RO SD SE SI SK SL SZ TR TZ UG ZM ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EC EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NI NO NZ OM PH PL PT RO RU SC SD SE SG SK SL TJ TM TN TR TT TZ UA UG US

UZ VC VN YU ZA ZM ZW

AU 2003238883 A1 20031222 (200445) AU 2003238883 A8 20031222 (200559)

# APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
WO 2003104212	A1	WO 2003-US17600	20030604
AU 2003238883	A1	AU 2003-238883	20030604
AU 2003238883	A8	AU 2003-238883	20030604

#### FILING DETAILS:

PATENT NO KIND \_\_\_\_\_\_ AU 2003238883 Al Based on WO 2003104212 AU 2003238883 A8 Based on WO 2003104212 PRIORITY APPLN. INFO: IN 2002-CH425 20020605 2004-071377 [07] WPIDS 2004-071376 [07] CR AB WO2003104212 A UPAB: 20050915 NOVELTY - Amorphous form of (2-(4-((4-chlorophenyl )-phenylmethyl)-1-piperazinyl)ethoxy)acetic acid dihydrochloride (cetirizine dihydrochloride) is new. DETAILED DESCRIPTION - An INDEPENDENT CLAIM is included for preparation of the amorphous form of cetirizine dihydrochloride. ACTIVITY - Antiallergic; Antiinflammatory; Ophthalmological; Antipruritic; Dermatological. MECHANISM OF ACTION - None given. USE - In a pharmaceutical formulations for treating allergic syndromes e.g. chronic and acute allergic rhinitis, allergic conjunctivitis, pruritus, or urticaria. ADVANTAGE - The amorphous form has moisture content of 0.3 - 12 (preferably 1.8 - 5.6) % by KF method. The amorphous form can be obtained by simple, eco-friendly and commercially viable scalable. Dwg.0/1 L18 ANSWER 12 OF 17 WPIDS COPYRIGHT 2006 THE THOMSON CORP on STN ACCESSION NUMBER: 2004-071376 [07] CROSS REFERENCE: 2004-071377 [07] WPTDS CROSS REFERENCE: DOC. NO. CPI: C2004-029507 TITLE: New crystalline form of cetirizine dihydrochloride useful for treating e.g. allergic rhinitis, allergic conjunctivitis and pruritis. DERWENT CLASS: B03 RAJAN, S T; REDDY, M S; SHANKAR, R R; VARDHAN, S V; INVENTOR(S): MANNE, S R; RANGA, R S; SRINIVASAN, T R; SUNKARA, V V (REDD-N) REDDY'S LAB LTD; (REDD-N) REDDYS LAB LTD PATENT ASSIGNEE(S): 103 COUNTRY COUNT: PATENT INFORMATION: PATENT NO KIND DATE WEEK LA PG \_\_\_\_\_ WO 2003104211 A2 20031218 (200407) \* EN 22 RW: AT BE BG CH CY CZ DE DK EA EE ES FI FR GB GH GM GR HU IE IT KE LS LU MC MW MZ NL OA PT RO SD SE SI SK SL SZ TR TZ UG ZM ZW W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EC EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NI NO NZ OM PH PL PT RO RU SC SD SE SG SK SL TJ TM TN TR TT TZ UA UG US UZ VC VN YU ZA ZM ZW AU 2003237394 A1 20031222 (200445) IN 2002000425 I4 20050304 (200555) AU 2003237394 A8 20031222 (200559) APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
WO 2003104211	A2	WO 2003-US17672	20030604
AU 2003237394	A1	AU 2003-237394	20030604
IN 2002000425	I4	IN 2002-CH425	20020605
AU 2003237394	A8	AU 2003-237394	20030604

## FILING DETAILS:

PATENT NO	KIND	PATENT NO
AU 2003237394	Al Based on	WO 2003104211
AU 2003237394	A8 Based on	WO 2003104211

PRIORITY APPLN. INFO: IN 2002-CH425 20020605

AN 2004-071376 [07] WPIDS

CR 2004-071377 [07]

AB W02003104211 A UPAB: 20050915

NOVELTY - A crystalline form of cetirizine

dihydrochloride (I) is new.

DETAILED DESCRIPTION - An INDEPENDENT CLAIM is included for the preparation of (I).

ACTIVITY - Antiallergic; Antipruritic; Dermatological; Ophthalmological.

MECHANISM OF ACTION - Histamine H1 receptor antagonist.

USE - For treating allergic syndromes (e.g. chronic and acute allergic rhinitis, allergic conjunctivitis, pruritis and urticaria).

ADVANTAGE - The crystalline form of cetirizine dihydrochloride is orally active and long acting histamine H1 receptor antagonist; exhibits less sedation, low anticholinergic activity and longer acting duration with improved patient compliance.

Dwg.0/3

WPIDS

L18 ANSWER 13 OF 17 WPIDS COPYRIGHT 2006 THE THOMSON CORP on STN

ACCESSION NUMBER:

2003-210201 [20]

DOC. NO. CPI:

C2003-053581

TITLE:

Tablet used for treating e.g.

disorders associated with allergic rhinitis, ocular pruritus and sneezing, comprises cetirizine and pseudoephedrine in distinct segments and alkalizing

agent.

DERWENT CLASS:

B05 B07

INVENTOR(S):
PATENT ASSIGNEE(S):

BERWAER, M; DELEERS, M; FANARA, D; GUICHAUX, A
(UNIO) UCB SA; (UNIO) UCB FARCHIM SA; (BERW-I)

BERWAER M; (DELE-I) DELEERS M; (FANA-I) FANARA D;

(GUIC-I) GUICHAUX A

COUNTRY COUNT:

101

PATENT INFORMATION:

PATENT NO	KIND	DATE	WEEK	LA	PG
WO 2003002098	A1 20	0030109	(200320)*	EN 1	3

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ NL OA PT SD SE SL SZ TR TZ UG ZM ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EC EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ OM PH PL PT RO RU SD SE SG SI SK SL TJ TM TN TR TT TZ UA UG US UZ

VN YU ZA ZM ZW EP 1404304 A1 20040407 (200425) EN R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT RO SE SI TR SK 2003001550 A3 20040504 (200433) KR 2004007756 A 20040124 (200435) AU 2002345024 A1 20030303 (200452) CZ 2003003454 A3 20040818 (200457) US 2004170690 A1 20040902 (200458)
HU 2004000386 A2 20040830 (200465)
BR 2002010650 A 20041005 (200475)
CN 1520285 A 20040811 (200476)
NZ 530289 A 20041269 (200479) JP 2004536829 W 20041209 (200481) ZA 2003009720 A 20050223 (200519) NO 2003005798 A 20040227 (200561) 43 28 MX 2003010430 A1 20050101 (200564) US 2006034928 A1 20060216 (200614) B2 20060321 (200621) US 7014867

## APPLICATION DETAILS:

PATENT 1	40 KI	ND	A:	PPLICATION	DATE
WO 20030	002098 A			2002-EP6342	
EP 14043	304 A	1	EP	2002-743173	20020610
			ŴΟ	2002-EP6342	20020610
SK 20030	001550 A	3	WO	2002-EP6342	20020610
				2003-1550	20020610
KR 20040	007756 A		KR	2003-717054	20031227
AU 20023	345024 A	1	AU	2002-345024	20020610
CZ 20030	003454 A	3	WO	2002-EP6342	20020610
			CZ	2003-3454	20020610
US 20043	170690 A	1 Provisional	US	2001-301250P	20010628
			WO	2002-EP6342	20020610
			US	2003-481264	20031219
HU 20040	000386 A	2	WO	2002-EP6342	20020610
			HU	2004-386	20020610
BR 20020	010650 A		BR	2002-10650	20020610
			WO	2002-EP6342	20020610
CN 15202	285 A			2002-812975	20020610
NZ 53028	89 A			2002-530289	20020610
				2002-EP6342	20020610
JP 20045	536829 W			2002-EP6342	20020610
				2003-508337	20020610
ZA 20030	· <del>-</del> -			2003-9720	20031215
NO 20030	005798 A			2002-EP6342	20020610
				2003-5798	20031223
MX 20030	010430 A	1		2002-EP6342	20020610
				2003-10430	20031114
US 20060	034928 A	1 Provisional		2001-301250P	
		Div ex		2002-EP6342	20020610
		Div ex		2003-481264	20031219
				2005-251895	20051018
US 70148	867 B	2		2002-EP6342	20020610
			US	2003-481264	20031219

#### FILING DETAILS:

PATE	NT NO	KIN	ID		I	PATENT NO
EP 1	404304	A1	Based	on	WO	2003002098
SK 2	003001550	Α3	Based	on	WO	2003002098
AU 2	002345024	Α1	Based	on	WO	2003002098
CZ 2	003003454	A3	Based	on	WO	2003002098
HU 2	004000386	A2	Based	on	WO	2003002098
BR 2	002010650	Α	Based	on	WO	2003002098
NZ 5	30289	Α	Based	on	WO	2003002098
JP 2	004536829	W	Based	on	WO	2003002098
MX 2	003010430	A1	Based	on	WO	2003002098
us 7	014867	В2	Based	on	WO	2003002098
RITY	APPLN. INFO:	US	2001-	-301250P	2	20010628; EI
		2.0	101_11	5007	2001	10628

PRIOR P 2001-115807 20010628

2003-210201 [20] WPIDS AN WO2003002098 A UPAB: 20030324 AB

NOVELTY - Tablet (T) comprises at least two distinct segments comprising cetirizine (cl) and pseudoephedrine (p). (T) Also comprises less than 5 weight% of an alkalizing agent relative to the total weight of (T).

ACTIVITY - Antiallergic; Antiinflammatory; Virucide; Antipyretic; Antipruritic; Ophthalmological.

MECHANISM OF ACTION - None given in the source material.

USE - Used for preventing or treating disorders or conditions associated with rhinitis, cold, flu, cold and flu-like symptoms, allergic rhinitis, relief of nasal congestion, seasonal rhinitis, sneezing, rhinorrhea, nasal, ocular pruritus, redness of the eyes, tearing or sneezing (all claimed). (T) Is also useful as an antiallergic, antihistaminic, bronchodilatory or antispasmodic agent.

Dwg. 0/0

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ACCESSION NUMBER: 2003356001 EMBASE

TITLE: Anaphylactic reactions to tolperisone (Mydocalm®).

Ribi C.; Vermeulen C.; Hauser C. AUTHOR:

Dr. C. Ribi, Allergy Unit, Division of Immunology and CORPORATE SOURCE:

Allergy, University Hospital Geneva, CH-1211 Geneva,

Switzerland. Camillo.Ribi@hcuge.ch

Swiss Medical Weekly, (28 Jun 2003) Vol. 133, No. SOURCE:

25-26, pp. 369-371. .

Refs: 6

ISSN: 1424-7860 CODEN: SMWWAI

Switzerland COUNTRY: DOCUMENT TYPE: Journal; Article

FILE SEGMENT: 026 Immunology, Serology and Transplantation

> Drug Literature Index 037 038 Adverse Reactions Titles

LANGUAGE: English SUMMARY LANGUAGE: English

Entered STN: 18 Sep 2003 ENTRY DATE:

Last Updated on STN: 18 Sep 2003

Four patients with anaphylaxis attributed to the intake of the AB centrally acting muscle relaxant tolperisone hydrochloride (Mydocalm®) were observed at the Emergency Department of the Geneva University Hospital between November 2001 and March 2003. patients were middle-aged women who took tolperisone for chronic

> Shears 571-272-2528 Searcher :

muscular pain. All reactions occurred within an hour after oral intake of this drug frequently prescribed in Switzerland. The severity of anaphylaxis ranged from urticarial reactions to shock with arterial hypotension. Prick-to-prick skin testing performed in one patient with a tablet of tolperisone diluted in water was negative. Its globally restricted commercialisation may explain the lack of reports on such adverse effects in the MedLine database. Anaphylactic reactions to this drug, however, are mentioned in other sources such as the Swiss Drug Compendium and the WHO drug reaction database. Together, these findings suggest that anaphylaxis to tolperisone is not uncommon and should be known to physicians in countries where this drug is available.

L18 ANSWER 15 OF 17 EMBASE COPYRIGHT (c) 2006 Elsevier B.V. All rights reserved on STN

ACCESSION NUMBER: 2003048852 EMBASE

TITLE: [Antihistamines in dermatology].

LEKI PRZECIWHISTAMINOWE W DERMATOLOGII.

AUTHOR: Czarnecka-Operacz M.; Silny W.

SOURCE: Przeglad Dermatologiczny, (2002) Vol. 89, No. 6, pp.

435-443. . Refs: 55

ISSN: 0033-2526 CODEN: PRDEA7

COUNTRY: Poland

DOCUMENT TYPE: Journal; Article

FILE SEGMENT: 013 Dermatology and Venereology

030 Pharmacology

037 Drug Literature Index

039 Pharmacy

LANGUAGE: Polish

SUMMARY LANGUAGE: English; Polish

ENTRY DATE: Entered STN: 7 Feb 2003

Last Updated on STN: 7 Feb 2003

AB Antihistamines are widely used in the treatment of various skin diseases. Both allergic and nonallergic inflammatory dermatoses are indications for antihistaminic therapy. For example acute and chronic urticaria, angioedema, atopic dermatitis, allergic and nonallergic eczema, airborne dermatitis, various drug induced reactions and drug eruptions, fotodermatoses, nodular prurigo, circumscribed neurodermitis and many other pruritic skin diseases are treated with antihistamines. Classical antihistamines are used in acute clinical cases in which parenteral route of application is necessary while second generation of antihistamines is prescribed in case of various inflammatory eosinophil or neutrophil related skin disorders. Therefore indications for antihistaminic therapy include also such diseases as bullous pemphigoid, pemphigus vulgaris, dermatitis herpetiformis, linear IgA bullous dermatosis etc. It seems that dermatology is the field of medical sciences in which antihistamines are applied most often and that is why we should be familiar with mechanisms of action of this group of drugs, all possible side effects and contrindications.

L18 ANSWER 16 OF 17 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on STN DUPLICATE 1

ACCESSION NUMBER: 2002:213411 BIOSIS DOCUMENT NUMBER: PREV200200213411

TITLE: Levocetirizine in allergic diseases: An open

multicenter practice study on efficacy and safety.

Original Title: Levocetirizin bei allergischen

Erkrankungen: Eine offene multizentrische Praxisstudie

zur Wirksamkeit und Vertraeglichkeit.

AUTHOR(S): Klimek, L. [Reprint author]; Hundorf, I.

Allergologie, Umweltmedizin, HNO, An den Quellen 10, CORPORATE SOURCE:

D-65183, Wiesbaden, Germany

Allergologie, (Januar, 2002) Vol. 25, No. 1, pp. S1-S7. SOURCE:

print.

CODEN: ALLRDI. ISSN: 0344-5062.

DOCUMENT TYPE:

Article

LANGUAGE:

German

ENTRY DATE:

Entered STN: 27 Mar 2002

Last Updated on STN: 27 Mar 2002

In an open multicenter practice study the clinical efficacy and safety AB of the selective, peripheral H1-receptor antagonist levocetirizine was studied systematically in 17,638 patients with allergic diseases of the airways and skin. The mean age of the patients was 38.1+-16.0 years. 14,319 patients suffered from allergic diseases of the respiratory airways, 4,704 from allergic diseases of the skin. During a mean observation period of 32 days the patients received 5 mg levocetirizine daily (1 film-coated tablet Xusal(R)). The changes of clinical symptoms were documented at the beginning and end of the therapy. Optional an interim examination could be performed. The efficacy of levocetirizine was assessed according to the severity of the nasal symptoms pruritus, rhinorrhea, sneezing, obstruction, and if applicable of asthmoid symptoms and ocular complaints. Skin symptoms like pruritus, wheal, flare and eczema were recorded. At the end of the treatment period a total symptom relief or clear improvement for allergic symptoms of the airways, eyes and skin was achieved on average in 80-90% of the patients. The global efficacy of levocetirizine was assessed with "very good" to "good" in 86.9% of all cases. In 82.8% of the patients a fast onset of action occurred within 60 minutes. In 95.5% of all cases the safety of levocetirizine was assessed with "very good" to "good". During the practice study 407 adverse events occurred in 299 (1.7%) of 17,638 patients in all. Out of these adverse events 369 were assessed as adverse drug reaction with mild to moderate impairment only. The remaining 38 reports showed no causal relationship with levocetirizine. 7 events were classified as serious but without possible causal relationship to levocetirizine. On the basis of this data from a large number of patients we state that 5 mg levocetirizine daily provides fast and powerful relief of symptoms in patients with allergic diseases of airways and skin with good tolerability at the same time.

L18 ANSWER 17 OF 17 EMBASE COPYRIGHT (c) 2006 Elsevier B.V. All rights reserved on STN

ACCESSION NUMBER:

2002048654 EMBASE

TITLE:

[Levocetirizine in allergic diseases - An

open multicenter practice study on efficacy and

safety].

LEVOCETIRIZIN BEI ALLERGISCHEN ERKRANKUNGEN: EINE

OFFENE MULTIZENTRISCHE PRAXISSTUDIE ZUR WIRKSAMKEIT UND

VERTRAGLICHKEIT.

AUTHOR:

Klimek L.; Hundorf I.

CORPORATE SOURCE:

Dr. L. Klimek, HNO, Allergologie, Umweltmedizin, An den

Quellen 10, D-65183 Wiesbaden, Germany

SOURCE:

Allergologie, (2002) Vol. 25, No. 1, pp. S1-S7. .

Shears 571-272-2528 Searcher :

Refs: 19

ISSN: 0344-5062 CODEN: ALLRDI

COUNTRY:

Germany

DOCUMENT TYPE:

Journal; Article

FILE SEGMENT:

011 Otorhinolaryngology

TIDE DECIDENT.

013 Dermatology and Venereology

026

Immunology, Serology and Transplantation

037 Drug Literature Index

LANGUAGE:

German

SUMMARY LANGUAGE:

English; German

ENTRY DATE:

Entered STN: 14 Feb 2002

Last Updated on STN: 14 Feb 2002

In an open multicenter practice study the clinical efficacy and safety AB of the selective, peripheral H(1)-receptor antagonist levocetirizine was studied systematically in 17,638 patients with allergic diseases of the airways and skin. The mean age of the patients was 38.1 ± 16.0 years. 14,319 patients suffered from allergic diseases of the respiratory airways, 4,704 from allergic diseases of the skin. During a mean observation period of 32 days the patients received 5 mg levocetirizine daily (1 film-coated tablet Xusal.RTM.). The changes of clinical symptoms were documented at the beginning and end of the therapy. Optional an interim examination could be performed. The efficacy of levocetirizine was assessed according to the severity of the nasal symptoms pruritus, rhinorrhea, sneezing, obstruction, and if applicable of asthmoid symptoms and ocular complaints. Skin symptoms like pruritus, wheal, flare and eczema were recorded. At the end of the treatment period a total symptom relief or clear improvement for allergic symptoms of the airways, eyes and skin was achieved on average in 80 - 90% of the patients. The global efficacy of levocetirizine was assessed with "very good" to "good" in 86.9% of all cases. In 82.8% of the patients a fast onset of action occurred within 60 minutes. In 95.5% of all cases the safety of levocetirizine was assessed with "very good" to "good". During the practice study 407 adverse events occurred in 299 (1.7%) of 17,638 patients in all. Out of these adverse events 369 were assessed as adverse drug reaction with mild to moderate impairment only. The remaining 38 reports showed no causal relationship with levocetirizine. 7 events were classified as serious but without possible causal relationship to levocetirizine. On the basis of this data from a large number of patients we state that 5 mg levocetirizine daily provides fast and powerful relief of symptoms in patients with allergic diseases of airways and skin with good tolerability at the same time.

FILE 'CAPLUS' ENTERED AT 16:23:03 ON 10 APR 2006

L19 8 SEA ABB=ON PLU=ON L6 AND (ORAL? OR PER OS OR MOUTH) (S) (AD MIN? OR DOSAGE OR DOSE OR DOSING OR INTAK?)

L20 0 SEA ABB=ON PLU=ON L19 NOT L9

L21 O SEA ABB=ON PLU=ON (L3 OR L5) AND KF

FILE 'MEDLINE, BIOSIS, EMBASE, WPIDS, CONFSCI, SCISEARCH, JICST-EPLUS, JAPIO' ENTERED AT 16:24:48 ON 10 APR 2006

L22 3 SEA ABB=ON PLU=ON L21

L23 0 SEA ABB=ON PLU=ON L22 NOT L17

(FILE 'CAPLUS, MEDLINE, BIOSIS, EMBASE, WPIDS, CONFSCI, SCISEARCH, JICST-EPLUS, JAPIO' ENTERED AT 16:26:11 ON 10 APR 2006)

L24 9592 SEA ABB=ON PLU=ON "REDDY M"?/AU

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-Author(s)
L25
          1718 SEA ABB=ON PLU=ON "RAJAN S"?/AU
          2316 SEA ABB=ON PLU=ON "RAO U"?/AU
L26
             5 SEA ABB=ON PLU=ON "RAMAYYA V"?/AU
L27
L28
             2 SEA ABB=ON PLU=ON L24 AND L25 AND L26 AND L27
            60 SEA ABB=ON PLU=ON L24 AND (L25 OR L26 OR L27)
L29
            12 SEA ABB=ON PLU=ON L25 AND (L27 OR L26)
L30
L31
             2 SEA ABB=ON PLU=ON L26 AND L27
         13557 SEA ABB=ON PLU=ON L24 OR L25 OR L26 OR L27
L32
             8 SEA ABB=ON PLU=ON (L24 OR L25 OR L26 OR L27 OR L29 OR
L33
               L32) AND (L3 OR L5)
            18 SEA ABB=ON PLU=ON L28 OR L30 OR L31 OR L33
L34
             9 DUP REM L34 (9 DUPLICATES REMOVED)
L35
L35 ANSWER 1 OF 9 CAPLUS COPYRIGHT 2006 ACS on STN DUPLICATE 1
ACCESSION NUMBER:
                        2005:611976 CAPLUS
DOCUMENT NUMBER:
                        143:139152
                        Preparation of polymorphs of ezetimibe
TITLE:
INVENTOR(S):
                        Sundaram, Venkataraman; Rajan, Srinivasan
                        Thirumalai; Ramayya, Vaddadi Pattabhi
                        ; Vardhan, Sunkara Vishnu; Subrahmanyam, Bulusu;
                        Sasikala, Cheemalapati Venkata Annapurna
                        Reddy's Laboratories Ltd., India; Reddy's
PATENT ASSIGNEE(S):
                        Laboratories, Inc.
SOURCE:
                        PCT Int. Appl., 26 pp.
                        CODEN: PIXXD2
DOCUMENT TYPE:
                        Patent
                        English
LANGUAGE:
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:
     DAMENIE NO
                                           ADDITCARTON NO
                                                                  שתעת
                        TENT
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PA.	rent :	NO.			KIN	D	DATE		1	APPL	ICAT:	ION !	ΝΟ.		D	ATE
WO	2005	0628	<b></b> 97		A2		2005	0714	1	WO 2	004-1	JS43	157		2	0041223
	W:	ΑE,	AG,	AL,	AM,	ΑT,	AU,	ΑZ,	BA,	BB,	BG,	BR,	BW,	BY,	BZ,	CA,
		CH,	CN,	CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	EG,	ES,	FI,
		GB,	GD,	GE,	GH,	GM,	HR,	ΗU,	ID,	IL,	IN,	IS,	JP,	ΚE,	KG,	KP,
		KR,	ΚZ,	LC,	LK,	LR,	LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,
		MX,	MZ,	NA,	NI,	NO,	NZ,	OM,	PG,	PH,	PL,	PT,	RO,	RU,	SC,	SD,
		SE,	SG,	SK,	SL,	SY,	ТJ,	TM,	TN,	TR,	TT,	ΤZ,	UA,	UG,	US,	UZ,
		VC,	VN,	YU,	ZA,	ZM,	ZW									
	RW:	B₩,	GH,	GM,	ΚE,	LS,	MW,	MZ,	NA,	SD,	SL,	SZ,	TZ,	ŪG,	ZM,	ZW,
		AM,	ΑZ,	BY,	KG,	ΚZ,	MD,	RU,	ТJ,	TM,	AT,	BE,	BG,	CH,	CY,	CZ,
		DE,	DK,	EE,	ES,	FI,	FR,	GB,	GR,	HU,	ΙE,	IS,	IT,	LT,	LU,	MC,
		NL,	PL,	PT,	RO,	SE,	SI,	SK,	TR,	ΒF,	ВJ,	CF,	CG,	CI,	CM,	GA,
		GN,	GQ,	GW,	ML,	MR,	ΝE,	SN,	TD,	ΤG						
US	2005	1710	80		A1		2005	0804	1	US 2	004-	2257	0		2	0041223
PRIORITY	Y APP	LN.	INFO	.:						IN 2	003-	CH10	49	i	A 2	0031223

AB The present invention relates to novel crystalline forms and amorphous form of ezetimibe and the processes for the preparation thereof. Thus, a crystalline

form of ezetimibe was dissolved in MeOH and the solvent was evaporated to dryness to give the amorphous form.

L35 ANSWER 2 OF 9 CAPLUS COPYRIGHT 2006 ACS on STN DUPLICATE 2

ACCESSION NUMBER: 2005:185394 CAPLUS

DOCUMENT NUMBER: 142:280230

TITLE: A process for preparation of

(benzisothiazolylpiperazinylethyl)indolone

derivative (ziprasidone hydrochloride), useful as

antipsychotic agent

Reddy, Manne Satyanarayana; Venkatraman, Sundaram; INVENTOR(S):

> Rajan, Srinivasan Thirumalai; Narsapur, Sharat Pandurang; Kharkar, Manoj Ramesh;

Devarkonda, Surya Narayana; Reddy, Yarraguntla Sesha; Srinivasulu, Rangineni; Shukla, Deepak K.;

Lakhekar, Pushkar B.; Rao, Uppala Venkata

Bhaskar; Venkatesh, Mummadi

Reddy's Laboratories Limited, India; Reddy's PATENT ASSIGNEE(S):

Laboratories, Inc.

U.S. Pat. Appl. Publ., 10 pp. SOURCE:

CODEN: USXXCO

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.		DATE
US 2005049295 PRIORITY APPLN. INFO.:	A1	20050303	US 2004-868506 IN 2003-MA488	- <b>-</b> A	20040614 20030612
			IN 2004-CH222	Α	20040312

GΙ

The invention relates to improved processes for the preparation of AΒ (benzisothiazolylpiperazinylethyl)indolone hydrochloride derivative (I.HCl), useful as antipsychotic agent (no biol. data). Compound I.HCl (ziprasidone hydrochloride) was prepared via reduction of (chloroacetyl) indole derivative II (X = 0), amination of the obtained (chloroethyl) indole derivative II (X is absent) by 3-(1-piperazinyl)-1,2benzisothiazole, and subsequent hydrochloride salt formation of the formed ziprasidone.

Ι

L35 ANSWER 3 OF 9 CAPLUS COPYRIGHT 2006 ACS on STN DUPLICATE 3

ACCESSION NUMBER: 2005:78236 CAPLUS

DOCUMENT NUMBER: 142:162672

TITLE: Crystalline cetirizine monohydrochloride

INVENTOR(S): Reddy, Manne Satyanarayana; Rajan, Srinivasan

Thirumalai; Rao, Uppala Venkata Bhaskara; Reddy, Konda Srinivasa

PATENT ASSIGNEE(S): Reddy's Laboratories Limited, India; Reddy's

Laboratories, Inc.

SOURCE: U.S. Pat. Appl. Publ., 11 pp.

CODEN: USXXCO

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.		DATE
				-	
US 2005020608	A1	20050127	US 2004-809192		20040325
PRIORITY APPLN. INFO.:			IN 2003-MA252	Α	20030325

AB A novel crystalline form of cetirizine monohydrochloride and processes for making the crystalline form as well as compns., pharmaceutical compns., and methods utilizing the crystalline form are described. A process for preparation

of a crystalline form of cetirizine monohydrochloride, comprises (1) providing a solid residue of crude cetirizine monohydrochloride; (2) contacting the crude residue with a ketone solvent to cause separation of a solid mass; and (3) isolating the solid mass thereby obtaining the crystalline form of cetirizine monohydrochloride. Tablets for the treatment of allergic syndromes were formulated containing crystalline cetirizine monohydrochloride 10, CaCO3 500, PVP 17, Avicel 15, mannitol 400, maltodextrin 15, aspartame 3, and aroma 20 mg each.

L35 ANSWER 4 OF 9 CAPLUS COPYRIGHT 2006 ACS on STN DUPLICATE 4

ACCESSION NUMBER: 2004:493694 CAPLUS

DOCUMENT NUMBER: 141:54360

TITLE: Polymorphic crystalline forms of dihydrochloride

salts of cetirizine and processes for their

preparation

INVENTOR(S): Reddy, Manne Satyanarayana; Srinivasan,

Thirumalai Rajan; Uppala, Venkata Bhaskara Rao;

Vaddadi, Pattabhi Ramayya; Joga, Rajender

PATENT ASSIGNEE(S): Reddy's Laboratories Limited, India; Reddy's

Laboratories, Inc.

SOURCE: PCT Int. Appl., 37 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2004050647	A2	20040617	WO 2003-US38494	20031204
WO 2004050647	A3	20040902		
WO 2004050647	C1	20050303		

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA,

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CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI,
             GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP,
             KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW,
             MX, MZ, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE,
             SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC,
             VN, YU, ZA, ZM, ZW
         RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM,
             AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE,
             DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO,
             SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML,
             MR, NE, SN, TD, TG
                                             CA 2003-2488114
                                20040617
                                                                    20031204
    CA 2488114
                          AΑ
    AU 2003297640
                                20040623
                                            AU 2003-297640
                          Α1
                                                                    20031204
                                            US 2003-729856
    US 2004186112
                          A1
                                20040923
                                                                    20031204
PRIORITY APPLN. INFO.:
                                             IN 2002-MA908
                                                                 A 20021204
                                            WO 2003-US38494
                                                                    20031204
```

AB Crystalline polymorphic forms of the levorotatory and dextrorotatory cetirizine dihydrochloride salts are prepared by dissolving the salts in an a ketone-containing solvent (e.g., aqueous acetone), cooling the solution, and

collecting the crystalline precipitate

L35 ANSWER 5 OF 9 CAPLUS COPYRIGHT 2006 ACS on STN DUPLICATE 5

ACCESSION NUMBER: 2005:2182 CAPLUS

DOCUMENT NUMBER: 142:93859

TITLE: Process for the preparation of an amorphous

crystal form of the antiallergic cetirizine

dihydrochloride

INVENTOR(S): Reddy, Manne Satyanarayana; Rajan, Srinivasan

Thirumalai; Rao, Uppala Venkata Bhaskara; Reddy, Konda Srinivasa

PATENT ASSIGNEE(S): Reddy's Laboratories Limited, India; Reddy's

Laboratories, Inc.

SOURCE: U.S. Pat. Appl. Publ., 11 pp.

CODEN: USXXCO

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
	<b>-</b>			
US 2004266787	A1	20041230	US 2004-809193	20040325
PRIORITY APPLN. INFO.:			IN 2003-MA253 A	20030325

AB An amorphous form of the antiallergic compound cetirizine dihydrochloride, prepared by the base-promoted hydrolysis of the corresponding amide of certizine, extraction, followed by HCl salification, is prepared as are pharmaceutical compns. utilizing this crystalline form.

L35 ANSWER 6 OF 9 CAPLUS COPYRIGHT 2006 ACS on STN DUPLICATE 6

ACCESSION NUMBER:

2004:2869 CAPLUS

DOCUMENT NUMBER:

140:47583

TITLE:

Amorphous levocetririzine dihydrochloride compositions for treatment of allergies

INVENTOR(S):

Reddy, Manne Satyanarayana; Rajan, Srinavasan Thirumalai; Rao, Uppala

Venkata Bhaskara; Ramayya, Vaddadi

Pattabhi

PATENT ASSIGNEE(S): Reddy's Laboratories Limited, India; Reddy's

Laboratories, Inc.

SOURCE: PCT Int. Appl., 29 pp.

CODEN: PIXXD2

DOCUMENT TYPE: LANGUAGE:

Patent English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PA'	TENT	NO.			KIN	D -	DATE				ICAT				D.	ATE
WO	2004	0008	23		A1		2003	1231	1						2	0030623
							ΑU,									
		-					DE,									
		GE,	GH,	GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	ΚE,	KG,	ΚP,	KR,	KZ,
		LC,	LK,	LR,	LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,
		NI,	NO,	NZ,	OM,	PG,	PH,	PL,	PT,	RO,	RU,	SC,	SD,	SE,	SG,	SK,
		SL,	ТJ,	TM,	TN,	TR,	TT,	TZ,	UA,	UG,	US,	UZ,	VC,	VN,	YU,	ZA,
		ZM,	ZW													
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		BY,	KG,	KΖ,	MD,	RU,	ТJ,	TM,	AT,	BE,	BG,	CH,	CY,	CZ,	DE,	DK,
		EE,	ES,	FI,	FR,	GB,	GR,	HU,	ΙE,	IT,	LU,	MC,	NL,	PT,	RO,	SE,
		SI,	SK,	TR,	BF,	ВJ,	CF,	CG,	CI,	CM,	GΑ,	GN,	GQ,	GW,	$\mathtt{ML},$	MR,
		ΝE,	SN,	TD,	TG											
	2489						2003	1231							_	0030623
AU	2003	2778	55		A1		2004	0106		AU 2	003-	2778.	55		2	0030623
US	2004	1327	43		A1		2004	0708	1	US 2	003-	6018	44		2	0030623
CN	1662	515			Α		2005	0831		CN 2	003-	8144	16		2	0030623
PRIORIT	Y APP	LN.	INFO	.:						IN 2	002-	MA47	2	•	A 2	0020621
									1	WO 2	003-	US19	777	1	W 2	0030623

AB A process for the preparation of the amorphous form of levocetirizine dihydrochloride is described. A pharmaceutical composition comprising a prophylactically or therapeutically effective amount

of an amorphous form of levocetirizine dihydrochloride and pharmaceutical excipients is provided. The amorphous form of levocetirizine dihydrochloride is suitable for pharmaceutical purposes in the treatment of allergies, including ailments such as chronic and acute allergic rhinitis, allergic conjunctivitis, pruritus, urticaria and the like.

REFERENCE COUNT:

THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L35 ANSWER 7 OF 9 CAPLUS COPYRIGHT 2006 ACS on STN DUPLICATE 7

ACCESSION NUMBER:

2003:991495 CAPLUS

DOCUMENT NUMBER:

140:47519

3

TITLE:

Process for the preparation of an amorphous form

of [2-[4-[(4-chlorophenyl

)phenylmethyl]-1- piperazinyl]ethoxy]
acetic acid dihydrochloride (cetirizine

dihydrochloride)

INVENTOR(S):

Reddy, Manne Satyanarayana; Rajan,

Srinivasan Thirumalai; Shankar, Ranga Ravi;

Vardhan, Sunkara Vishnu

PATENT ASSIGNEE(S): Dr.Reddy's Laboratories Ltd., India; Dr.Reddy's

Laboratories, Inc.

SOURCE: PCT Int. Appl., 21 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

PA	TENT :	NO.			KIN	D	DATE		1	APPL	ICAT:	ION I	.00		D	ATE	
WO	2003	1042	 12		A1	_	2003	1218	,	WO 2	003-1	US17	600		2		
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		CN,	co,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	ES,	FI,	GB,	GD,	
		GE,	GH,	GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	ΚE,	KG,	ΚP,	KR,	ΚZ,	
		LC,	LK,	LR,	LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	ΜZ,	
		NI,	NO,	NZ,	OM,	PH,	PL,	PT,	RO,	RU,	SC,	SD,	SE,	SG,	SK,	SL,	
		ТJ,	TM,	TN,	TR,	TT,	TZ,	UA,	UG,	US,	UZ,	VC,	VN,	YU,	ZA,	ZM,	ZW
	RW:	GH,	GM,	KE,	LS,	MW,	MZ,	SD,	SL,	SZ,	TZ,	UG,	ZM,	ZW,	AM,	ΑZ,	
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		EE,	ES,	FI,	FR,	GB,	GR,	HU,	ΙE,	IT,	LU,	MC,	NL,	PT,	RO,	SE,	
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AU	2003	2388	83		A1		2003	1222	3	AU 2	003-	2388	83		2	0030	604
PRIORIT	Y APP	LN.	INFO	.:						IN 2	002-1	MA42	5	j	A 2	0020	605
									1	WO 2	003-	US17	600	1	w 2	0030	604

AB A novel, amorphous form of [2-[4-[(4-Chlorophenyl))phenylmethyl]-1-piperazinyl]ethoxy]acetic acid

dihydrochloride, suitable for pharmaceutical formulations, is prepared

and X-ray diffraction patterns for it are presented.

REFERENCE COUNT: 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR

THIS RECORD. ALL CITATIONS AVAILABLE IN THE

RE FORMAT

L35 ANSWER 8 OF 9 CAPLUS COPYRIGHT 2006 ACS on STN DUPLICATE 8

ACCESSION NUMBER: 2003:991494 CAPLUS

DOCUMENT NUMBER: 140:42205

TITLE: Preparation of crystalline [2-[4-[(4-

chlorophenyl)phenylmethyl]-1piperazinyl]ethoxy]acetic acid

dihydrochloride (cetirizine dihydrochloride)

INVENTOR(S): Reddy, Manne Satyanarayana; Rajan,

Srinivasan Thirumalai; Shankar, Ranga Ravi;

Vardhan, Sunkara Vishnu

PATENT ASSIGNEE(S): Reddy's Laboratories Limited, India; Reddy's

Laboratories, Inc.

SOURCE: PCT Int. Appl., 22 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2003104211	A2	20031218	WO 2003-US17672	20030604
WO 2003104211	A3	20041223		

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W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH,
             CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD,
             GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ,
             LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ,
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             TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM,
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             NE, SN, TD, TG
                                20031222
                                           AU 2003-237394
                                                                   20030604
     AU 2003237394
                     A1
PRIORITY APPLN. INFO.:
                                           IN 2002-MA425
                                                               A 20020605
                                           WO 2003-US17672
                                                               W 20030604
OTHER SOURCE(S):
                         CASREACT 140:42205
     A crystalline form of cetirizine dihydrochloride (I), prepared by the
     salification of cetirizine with isopropanolic hydrogen chloride,
     having a defined X-ray diffraction pattern is presented, and
     pharmaceutical compns. containing I are presented.
L35 ANSWER 9 OF 9 CAPLUS COPYRIGHT 2006 ACS on STN DUPLICATE 9
ACCESSION NUMBER:
                        2002:977791 CAPLUS
DOCUMENT NUMBER:
                         138:61311
                         Novel crystalline forms of 4-[4-[4-
TITLE:
                         (hydroxydiphenylmethyl)-1-piperidinyl]-1-
                         hydroxybutyl]-\alpha, \alpha-
                         dimethylbenzeneacetic acid and its hydrochloride
INVENTOR(S):
                         Reddy, M. Satyanarayana; Rajan, S.
                         Thirumalai; Rao, U. V. Bhaskara
                         Reddy's Laboratories Ltd., India; Cord, Janet I.
PATENT ASSIGNEE(S):
                         PCT Int. Appl., 28 pp.
SOURCE:
                         CODEN: PIXXD2
DOCUMENT TYPE:
                         Patent
LANGUAGE:
                         English
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:
     PATENT NO.
                        KIND
                               DATE
                                           APPLICATION NO.
                                                                   DATE
                               -----
                                           _____
                                                                   _____
                        A2
                                20021227
                                           WO 2001-US23994
                                                                   20010731
     WO 2002102777
                        A3
     WO 2002102777
                               20030227
                               20031030
     WO 2002102777
                         C1
            AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH,
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             NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR,
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             CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
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Searcher : Shears 571-272-2528

R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC,

AA 20021227

PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR

Α

A2

20040216

20040324

CA 2450858

EP 1399422

EE 200400010

CA 2001-2450858

EP 2001-956057

EE 2004-10

20010731

20010731

20010731

Α	20040727	BR	2001-17054		20010731
Α	20040804	CN	2001-823379		20010731
T2	20050317	JP	2003-505320		20010731
C2	20060210	RU	2004-101045		20010731
A1	20040422	US	2003-362339		20031112
Α	20040914	ZA	2003-9557		20031209
Α	20041230	BG	2003-108435		20031211
		IN	2001-MA484	Α	20010618
		WO	2001-US23994	W	20010731
	A T2 C2 A1 A	A 20040804 T2 20050317 C2 20060210 A1 20040422 A 20040914	A 20040804 CN T2 20050317 JP C2 20060210 RU A1 20040422 US A 20040914 ZA A 20041230 BG IN	A 20040804 CN 2001-823379 T2 20050317 JP 2003-505320 C2 20060210 RU 2004-101045 A1 20040422 US 2003-362339 A 20040914 ZA 2003-9557 A 20041230 BG 2003-108435 IN 2001-MA484	A 20040804 CN 2001-823379 T2 20050317 JP 2003-505320 C2 20060210 RU 2004-101045 A1 20040422 US 2003-362339 A 20040914 ZA 2003-9557 A 20041230 BG 2003-108435 IN 2001-MA484 A

AB The present invention is related to novel polymorph of the title compound (fexofenadine) and fexofenadine-HCl and a process of preparation thereof. The present invention is also directed to provide pure novel polymorphs of fexofenadine and its hydrochloride by a simple process which is com. viable and environment friendly. Fexofenadine was prepared by the hydrolysis of a mixture of Me 4-[4-[4-(hydroxydiphenylmethyl)-1-piperidinyl]-1-oxobutyl]-α,α-dimethylbenzeneacetate-HCl and its isomer in MeOH with aqueous NaOH solution After completion of the reaction, NaBH4 was added to reduce the carbonyl group to give the crude fexofenadine. The compound was purified by repeated crystallization in MeOH and converted to the polymorph

by treatment with boling toluene.

FILE 'HOME' ENTERED AT 16:29:33 ON 10 APR 2006

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(FILE 'REGISTRY' ENTERED AT 15:55:10 ON 10 APR 2006)
                DEL HIS Y
                D COST
     FILE 'REGISTRY' ENTERED AT 16:10:11 ON 10 APR 2006
                E LEVOCETIRIZINE/CN 5
              2 SEA ABB=ON PLU=ON (LEVOCETIRIZINE/CN OR "LEVOCETIRIZINE
L1
                DIHYDROCHLORIDE"/CN)
     FILE 'REGISTRY' ENTERED AT 16:10:27 ON 10 APR 2006
                D 1-2 IDE
                E CETIRIZINE DIHYDROCHLORIDE/CN
              2 SEA ABB=ON PLU=ON (CETIRIZINE/CN OR "CETIRIZINE DIHYDROCH
L2
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                D IDE
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L3
           1502 SEA ABB=ON PLU=ON (CHLOROPHENYL? OR (CL OR CHLORO) (W) (PH
L4
                OR PHENYL?))(S)ACETIC
                D KWIC
L5
             31 SEA ABB=ON PLU=ON L4(S)PIPERAZIN?
             80 SEA ABB=ON PLU=ON (L3 OR L5) AND (THERAP? OR TREAT? OR
L6
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              3 SEA ABB=ON PLU=ON I.6 AND (EXCIPIENT OR (STABILIS? OR
T.7
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             19 SEA ABB=ON PLU=ON L6 AND (ORAL? OR TABLET OR PILL OR
1.8
                CAPSUL? OR PER OS OR MOUTH)
             21 S L6 AND ADMIN?
L*** DEL
L*** DEL
             32 S L7 OR L8 OR L9
                D KWIC L5
                D KWIC L9
                D KWIC L8
             19 SEA ABB=ON PLU=ON L7 OR L8
L9
                D OUE L7
                D QUE L8
                D L9 1-19 .BEVSTR
     FILE 'MEDLINE, BIOSIS, EMBASE, WPIDS, CONFSCI, SCISEARCH,
     JICST-EPLUS, JAPIO' ENTERED AT 16:16:58 ON 10 APR 2006
            416 SEA ABB=ON PLU=ON L6
L10
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L11
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L12
              4 SEA ABB=ON PLU=ON L12 AND (CRYSTAL? OR CRYST## OR
L13
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L14
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L15
                SOLID?)
             17 SEA ABB=ON PLU=ON L10 AND (TABLET OR PILL OR CAPSUL? OR
L16
                SOLID?)
             18 SEA ABB=ON PLU=ON L11 OR L13 OR L15 OR L16
L17
L18
             17 DUP REM L17 (1 DUPLICATE REMOVED)
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Searcher : Shears 571-272-2528

D 1-17 IBIB ABS

L19 L20 L21	8		L6 AND (ORAL? OR PER OS OR MOUTH) (S) (AD OSE OR DOSING OR INTAK?) L19 NOT L9
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L24		SEA ABB=ON PLU=ON	
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		SEA ABB=ON PLU=ON	
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L30			L25 AND (L27 OR L26)
L31	2	SEA ABB=ON PLU=ON	L26 AND L27
		SEA ABB=ON PLU=ON	
L33			(L24 OR L25 OR L26 OR L27 OR L29 OR
		L32) AND (L3 OR L5)	
L34	18		L28 OR L30 OR L31 OR L33
L35	9	DUP REM L34 (9 DUPL	ICATES REMOVED)
		D 1-9 IBIB ABS	

FILE 'HOME' ENTERED AT 16:29:33 ON 10 APR 2006

# FILE REGISTRY

Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 9 APR 2006 HIGHEST RN 879846-78-3 DICTIONARY FILE UPDATES: 9 APR 2006 HIGHEST RN 879846-78-3

New CAS Information Use Policies, enter HELP USAGETERMS for details.

TSCA INFORMATION NOW CURRENT THROUGH January 6, 2006

Please note that search-term pricing does apply when conducting SmartSELECT searches.

Structure search iteration limits have been increased. See HELP SLIMI for details.

REGISTRY includes numerically searchable data for experimental and predicted properties as well as tags indicating availability of

experimental property data in the original document. For information on property searching in REGISTRY, refer to:

http://www.cas.org/ONLINE/UG/regprops.html

FILE CAPLUS

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FILE COVERS 1907 - 10 Apr 2006 VOL 144 ISS 16 FILE LAST UPDATED: 9 Apr 2006 (20060409/ED)

Effective October 17, 2005, revised CAS Information Use Policies apply They are available for your review at:

http://www.cas.org/infopolicy.html

# FILE MEDLINE

FILE LAST UPDATED: 8 APR 2006 (20060408/UP). FILE COVERS 1950 TO DAT

On December 11, 2005, the 2006 MeSH terms were loaded.

The MEDLINE reload for 2006 is now (26 Feb.) available. For details on the 2006 reload, enter HELP RLOAD at an arrow prompt (=>). See also:

http://www.nlm.nih.gov/mesh/

http://www.nlm.nih.gov/pubs/techbull/nd04/nd04\_mesh.html

http://www.nlm.nih.gov/pubs/techbull/nd05/nd05\_med\_data\_changes.ht

http://www.nlm.nih.gov/pubs/techbull/nd05/nd05\_2006\_MeSH.html

OLDMEDLINE is covered back to 1950.

MEDLINE thesauri in the /CN, /CT, and /MN fields incorporate the MeSH 2006 vocabulary.

This file contains CAS Registry Numbers for easy and accurate substance identification.

# FILE BIOSIS

FILE COVERS 1969 TO DATE.

CAS REGISTRY NUMBERS AND CHEMICAL NAMES (CNs) PRESENT FROM JANUARY 1969 TO DATE.

RECORDS LAST ADDED: 5 April 2006 (20060405/ED)

### FILE EMBASE

FILE COVERS 1974 TO 10 Apr 2006 (20060410/ED)

EMBASE has been reloaded. Enter HELP RLOAD for details.

EMBASE is now updated daily. SDI frequency remains weekly (default) and biweekly.

This file contains CAS Registry Numbers for easy and accurate substance identification.

FILE WPIDS

FILE LAST UPDATED: 10 APR 2006 <20060410/UP>
MOST RECENT DERWENT UPDATE: 200624 <200624/DW>
DERWENT WORLD PATENTS INDEX SUBSCRIBER FILE, COVERS 1963 TO DATE

>>> FOR A COPY OF THE DERWENT WORLD PATENTS INDEX STN USER GUIDE, PLEASE VISIT:

http://www.stn-international.de/training\_center/patents/stn\_guide.pdf

>>> FOR DETAILS OF THE PATENTS COVERED IN CURRENT UPDATES, SEE http://scientific.thomson.com/support/patents/coverage/latestupdates/

>>> PLEASE BE AWARE OF THE NEW IPC REFORM IN 2006, SEE http://www.stn-international.de/stndatabases/details/ipc\_reform.html a http://scientific.thomson.com/media/scpdf/ipcrdwpi.pdf <<<

>>> UPCOMING NEW DWPI: EFFECTS ON SCRIPT RUNS - SEE NEWS MESSAGE <<<

FILE CONFSCI

FILE COVERS 1973 TO 24 Mar 2006 (20060324/ED)

CSA has suspended updates until further notice.

FILE SCISEARCH

FILE COVERS 1974 TO 7 Apr 2006 (20060407/ED)

SCISEARCH has been reloaded, see HELP RLOAD for details.

FILE JICST-EPLUS

FILE COVERS 1985 TO 3 APR 2006 (20060403/ED)

THE JICST-EPLUS FILE HAS BEEN RELOADED TO REFLECT THE 1999 CONTROLLED TERM (/CT) THESAURUS RELOAD.

FILE JAPIO

FILE LAST UPDATED: 3 APR 2006 <20060403/UP>
FILE COVERS APRIL 1973 TO DECEMBER 22, 2005

>>> GRAPHIC IMAGES AVAILABLE <<<

>>> NEW IPC8 DATA AND FUNCTIONALITY NOT YET AVAILABLE IN THIS FILE.

USE IPC7 FORMAT FOR SEARCHING THE IPC. WATCH THIS SPACE FOR FURTHE

DEVELOPMENTS AND SEE OUR NEWS SECTION FOR FURTHER INFORMATION

ABOUT THE IPC REFORM <<<

FILE HOME